

## WEST Search History

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DATE: Tuesday, May 11, 2004

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		<i>DB=PGPB,USPT,USOC,EPAB,JPAB,DWPI; THES=ASSIGNEE; PLUR=YES; OP=ADJ</i>	
<input type="checkbox"/>	L7	L6 and cellulosum	40
<input type="checkbox"/>	L6	L5 and (gene or dna or nucleic acid)	124
<input type="checkbox"/>	L5	L1 and epothilone b	159
<input type="checkbox"/>	L4	L3 and epothilone b	57
<input type="checkbox"/>	L3	L1 and biosynthesis	58
<input type="checkbox"/>	L2	L1 biosynthesis	0
<input type="checkbox"/>	L1	epothilone d or desoxyepothilone or desoxy adj5 epothilone	179

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**Search Results - Record(s) 1 through 20 of 57 returned.**☐ 1. Document ID: US 20040082026 A1**Using default format because multiple data bases are involved.**

L4: Entry 1 of 57

File: PGPB

Apr 29, 2004

PGPUB-DOCUMENT-NUMBER: 20040082026

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20040082026 A1

TITLE: Interferon alpha: remodeling and glycoconjugation of interferon alpha

PUBLICATION-DATE: April 29, 2004

## INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
DeFrees, Shawn	North Wales	PA	US	
Zopf, David	Wayne	PA	US	
Bayer, Robert	San Diego	CA	US	
Bowe, Caryn	Doylestown	PA	US	
Hakes, David	Willow Grove	PA	US	
Chen, Xi	Lansdale	PA	US	

US-CL-CURRENT: [435/68.1](#); [530/351](#)

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KWIC	Draw. Data
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☐ 2. Document ID: US 20040077836 A1

L4: Entry 2 of 57

File: PGPB

Apr 22, 2004

PGPUB-DOCUMENT-NUMBER: 20040077836

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20040077836 A1

TITLE: Granulocyte colony stimulating factor: remodeling and glycoconjugation of G-CSF

PUBLICATION-DATE: April 22, 2004

## INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
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DeFrees, Shawn	North Wales	PA	US
Zopf, David	Wayne	PA	US
Bayer, Robert	San Diego	CA	US
Bowe, Caryn	Doylestown	PA	US
Hakes, David	Willow Grove	PA	US
Chen, Xi	Lansdale	PA	US

US-CL-CURRENT: 530/351; 435/68.1

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KWIC	Draw D
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☐ 3. Document ID: US 20040063911 A1

L4: Entry 3 of 57

File: PGPB

Apr 1, 2004

PGPUB-DOCUMENT-NUMBER: 20040063911

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20040063911 A1

TITLE: Protein remodeling methods and proteins/peptides produced by the methods

PUBLICATION-DATE: April 1, 2004

## INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
DeFrees, Shawn	North Wales	PA	US	
Zopf, David	Wayne	PA	US	
Bayer, Robert	San Diego	CA	US	
Hakes, David	Willow Grove	PA	US	
Chen, Xi	Lansdale	PA	US	

US-CL-CURRENT: 530/351; 435/68.1, 530/395

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KWIC	Draw D
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☐ 4. Document ID: US 20040043446 A1

L4: Entry 4 of 57

File: PGPB

Mar 4, 2004

PGPUB-DOCUMENT-NUMBER: 20040043446

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20040043446 A1

TITLE: Alpha galactosidase a: remodeling and glycoconjugation of alpha galactosidase A

PUBLICATION-DATE: March 4, 2004

## INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
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DeFrees, Shawn	North Wales	PA	US
Zopf, David	Wayne	PA	US
Bayer, Robert	San Diego	CA	US
Bowe, Caryn	Doylestown	PA	US
Hakes, David	Willow Grove	PA	US
Chen, Xi	Lansdale	PA	US

US-CL-CURRENT: 435/68.1; 435/193, 435/208

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KMC	Draw. De
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☐ 5. Document ID: US 20040024216 A1

L4: Entry 5 of 57

File: PGPB

Feb 5, 2004

PGPUB-DOCUMENT-NUMBER: 20040024216

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20040024216 A1

TITLE: HMG-CoA reductase inhibitors and method

PUBLICATION-DATE: February 5, 2004

## INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Robl, Jeffrey A.	Newtown	PA	US	
Chen, Bang-Chi	Plainsboro	NJ	US	
Sun, Chong-Qing	East Windsor	NJ	US	

US-CL-CURRENT: 546/15; 546/79

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KMC	Draw. De
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☐ 6. Document ID: US 20040018598 A1

L4: Entry 6 of 57

File: PGPB

Jan 29, 2004

PGPUB-DOCUMENT-NUMBER: 20040018598

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20040018598 A1

TITLE: Bio-intermediates for use in the chemical synthesis of polyketides

PUBLICATION-DATE: January 29, 2004

## INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Santi, Daniel	San Francisco	CA	US	
Ashley, Gary	Alameda	CA	US	

Myles, David C.                      Kensington                      CA                      US

US-CL-CURRENT: 435/75; 435/118, 548/180

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KMC	Draw. De
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☐ 7. Document ID: US 20040014183 A1

L4: Entry 7 of 57

File: PGPB

Jan 22, 2004

PGPUB-DOCUMENT-NUMBER: 20040014183

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20040014183 A1

TITLE: Secondary metabolite congener distribution modulation

PUBLICATION-DATE: January 22, 2004

INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Licari, Peter J.	Fremont	CA	US	
Julien, Bryan	Oakland	CA	US	
Frykman, Scott	Hayward	CA	US	
Tsuruta, Hiroko	Emeryville	CA	US	

US-CL-CURRENT: 435/118; 435/252.3

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KMC	Draw. De
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☐ 8. Document ID: US 20030176473 A1

L4: Entry 8 of 57

File: PGPB

Sep 18, 2003

PGPUB-DOCUMENT-NUMBER: 20030176473

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20030176473 A1

TITLE: Derivatives of epothilone B and D and synthesis thereof

PUBLICATION-DATE: September 18, 2003

INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Taylor, Richard E.	South Bend	IN	US	
Chen, Yue	Mishawaka	IN	US	

US-CL-CURRENT: 514/365; 548/181, 548/204

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KMC	Draw. De
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☐ 9. Document ID: US 20030134860 A1

L4: Entry 9 of 57

File: PGPB

Jul 17, 2003

PGPUB-DOCUMENT-NUMBER: 20030134860  
PGPUB-FILING-TYPE: new  
DOCUMENT-IDENTIFIER: US 20030134860 A1

TITLE: Inhibitors of prenyl-protein transferase

PUBLICATION-DATE: July 17, 2003

## INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Dinsmore, Christopher J.	Schwenksville	PA	US	

US-CL-CURRENT: 514/250; 540/472

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KWIC	Draw. De
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☐ 10. Document ID: US 20030114450 A1

L4: Entry 10 of 57

File: PGPB

Jun 19, 2003

PGPUB-DOCUMENT-NUMBER: 20030114450  
PGPUB-FILING-TYPE: new  
DOCUMENT-IDENTIFIER: US 20030114450 A1

TITLE: Benzoquinone ansamycins

PUBLICATION-DATE: June 19, 2003

## INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Santi, Daniel	San Francisco	CA	US	
Myles, David C.	Kensington	CA	US	
Tian, Zong-Qiang	Fremont	CA	US	
Hutchinson, C. Richard	San Mateo	CA	US	
Johnson, Robert	Lafayette	CA	US	
Zhou, Yi-Qing	Lafayette	CA	US	
Feng, Li	Fremont	CA	US	

US-CL-CURRENT: 514/234.5; 514/252.13, 514/320, 514/337, 514/397, 540/456

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KWIC	Draw. De
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☐ 11. Document ID: US 20030096381 A1

L4: Entry 11 of 57

File: PGPB

May 22, 2003

PGPUB-DOCUMENT-NUMBER: 20030096381  
PGPUB-FILING-TYPE: new  
DOCUMENT-IDENTIFIER: US 20030096381 A1

TITLE: Production of polyketides

PUBLICATION-DATE: May 22, 2003

INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Julien, Bryan	Oakland	CA	US	
Katz, Leonard	Hayward	CA	US	
Khosla, Chaitan	Palo Alto	CA	US	

US-CL-CURRENT: [435/118](#); [435/193](#), [435/252.3](#), [435/320.1](#), [435/69.1](#), [536/23.2](#)

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KMOC	Draw. De
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☐ 12. Document ID: US 20030073205 A1

L4: Entry 12 of 57

File: PGPB

Apr 17, 2003

PGPUB-DOCUMENT-NUMBER: 20030073205  
PGPUB-FILING-TYPE: new  
DOCUMENT-IDENTIFIER: US 20030073205 A1

TITLE: Production of polyketides

PUBLICATION-DATE: April 17, 2003

INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Arslanian, Robert L.	Pacifica	CA	US	
Ashley, Gary	Alameda	CA	US	
Frykman, Scott	Hayward	CA	US	
Julien, Bryan	Oakland	CA	US	
Katz, Leonard	Oakland	CA	US	
Khosla, Chaitan	Palo Alto	CA	US	
Lau, Janice	San Mateo	CA	US	
Licari, Peter J.	Fremont	CA	US	
Regentin, Rika	Hayward	CA	US	
Santi, Daniel	San Francisco	CA	US	
Tang, Li	Foster City	CA	US	

US-CL-CURRENT: [435/117](#); [435/123](#), [435/252.3](#), [544/333](#), [546/148](#), [546/167](#), [546/281.7](#),  
[548/204](#)

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KMOC	Draw. De
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☐ 13. Document ID: US 20020193283 A1

L4: Entry 13 of 57

File: PGPB

Dec 19, 2002

PGPUB-DOCUMENT-NUMBER: 20020193283  
PGPUB-FILING-TYPE: new  
DOCUMENT-IDENTIFIER: US 20020193283 A1

TITLE: Inhibitors of prenyl-protein transferase

PUBLICATION-DATE: December 19, 2002

## INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Dinsmore, Christopher J.	Schwenksville	PA	US	
Bergman, Jeffrey M.	Perkasie	PA	US	

US-CL-CURRENT: 514/1

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KMC	Draw. De
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☐ 14. Document ID: US 20020137152 A1

L4: Entry 14 of 57

File: PGPB

Sep 26, 2002

PGPUB-DOCUMENT-NUMBER: 20020137152  
PGPUB-FILING-TYPE: new  
DOCUMENT-IDENTIFIER: US 20020137152 A1

TITLE: Fermentation process for epothilones

PUBLICATION-DATE: September 26, 2002

## INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Santi, Daniel	San Francisco	CA	US	
Metcalf, Brian	Moraga	CA	US	
Ashley, Gary	Alameda	CA	US	

US-CL-CURRENT: 435/118; 435/184, 435/252.3

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KMC	Draw. De
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☐ 15. Document ID: US 20020123497 A1

L4: Entry 15 of 57

File: PGPB

Sep 5, 2002

PGPUB-DOCUMENT-NUMBER: 20020123497  
PGPUB-FILING-TYPE: new  
DOCUMENT-IDENTIFIER: US 20020123497 A1



TITLE: Inhibitors of prenyl-protein transferase

PUBLICATION-DATE: September 5, 2002

INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Nguyen, Diem N.	North Wales	PA	US	
Stump, Craig A.	Pottstown	PA	US	
Williams, Theresa M.	Harleysville	PA	US	

US-CL-CURRENT: 514/249; 540/460

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KWIC	Draw. De
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☐ 16. Document ID: US 20020099007 A1

L4: Entry 16 of 57

File: PGPB

Jul 25, 2002

PGPUB-DOCUMENT-NUMBER: 20020099007

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20020099007 A1

TITLE: Inhibitors of prenyl-protein transferase

PUBLICATION-DATE: July 25, 2002

INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
deSolms, S. Jane	Collegeville	PA	US	
Stokker, Gerald E.	Gwynedd Valley	PA	US	
Shaw, Anthony W.	Lansdale	PA	US	

US-CL-CURRENT: 514/9; 530/317, 540/460

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KWIC	Draw. De
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☐ 17. Document ID: US 20020094977 A1

L4: Entry 17 of 57

File: PGPB

Jul 18, 2002

PGPUB-DOCUMENT-NUMBER: 20020094977

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20020094977 A1

TITLE: HMG-CoA reductase inhibitors and method

PUBLICATION-DATE: July 18, 2002

INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
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Robl, Jeffrey A.	Newtown	PA	US
Chen, Bang-Chi	Plainsboro	NJ	US
Sun, Chong-Qing	East Windsor	NJ	US

US-CL-CURRENT: [514/215](#); [514/291](#), [540/586](#), [546/80](#)

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KWC	Draw. De
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☐ 18. Document ID: US 20020068747 A1

L4: Entry 18 of 57

File: PGPB

Jun 6, 2002

PGPUB-DOCUMENT-NUMBER: 20020068747  
PGPUB-FILING-TYPE: new  
DOCUMENT-IDENTIFIER: US 20020068747 A1

TITLE: Inhibitors of prenyl-protein transferase

PUBLICATION-DATE: June 6, 2002

## INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Stump, Craig A.	Pottstown	PA	US	
Williams, Theresa M.	Harleysville	PA	US	

US-CL-CURRENT: [514/291](#); [514/393](#), [540/471](#)

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KWC	Draw. De
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☐ 19. Document ID: US 20020061901 A1

L4: Entry 19 of 57

File: PGPB

May 23, 2002

PGPUB-DOCUMENT-NUMBER: 20020061901  
PGPUB-FILING-TYPE: new  
DOCUMENT-IDENTIFIER: US 20020061901 A1

TITLE: HMG-CoA reductase inhibitors and method

PUBLICATION-DATE: May 23, 2002

## INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Robl, Jeffrey A.	Newtown	PA	US	
Chen, Bang-Chi	Plainsboro	NJ	US	
Sun, Chong-Qing	East Windsor	NJ	US	

US-CL-CURRENT: [514/290](#); [514/278](#), [546/15](#), [546/79](#), [546/93](#)

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KWIC	Draw D
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☐ 20. Document ID: US 20020052380 A1

L4: Entry 20 of 57

File: PGPB

May 2, 2002

PGPUB-DOCUMENT-NUMBER: 20020052380

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20020052380 A1

TITLE: Inhibitors of prenyl-protein transferase

PUBLICATION-DATE: May 2, 2002

INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Dinsmore, Christopher J.	Schwenksville	PA	US	
Bergman, Jeffrey M.	Perkasie	PA	US	

US-CL-CURRENT: 514/254.05; 514/218, 544/370

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KWIC	Draw D
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L3 and epothilone b

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☐ 21. Document ID: US 20020052363 A1

**Using default format because multiple data bases are involved.**

L4: Entry 21 of 57

File: PGPB

May 2, 2002

PGPUB-DOCUMENT-NUMBER: 20020052363  
PGPUB-FILING-TYPE: new  
DOCUMENT-IDENTIFIER: US 20020052363 A1

TITLE: Inhibitors of prenyl-protein transferase

PUBLICATION-DATE: May 2, 2002

INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Dinsmore, Christopher J.	Schwenksville	PA	US	
Bergman, Jeffrey M.	Perkasie	PA	US	

US-CL-CURRENT: [514/218](#); [514/254.05](#), [540/575](#), [544/370](#)

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KMC	Draw. Data
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☐ 22. Document ID: US 20020052028 A1

L4: Entry 22 of 57

File: PGPB

May 2, 2002

PGPUB-DOCUMENT-NUMBER: 20020052028  
PGPUB-FILING-TYPE: new  
DOCUMENT-IDENTIFIER: US 20020052028 A1

TITLE: Bio-intermediates for use in the chemical synthesis of polyketides

PUBLICATION-DATE: May 2, 2002

INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Santi, Daniel	San Francisco	CA	US	
Ashley, Gary	Alameda	CA	US	
Myles, David C.	Kensington	CA	US	

US-CL-CURRENT: [435/76](#); [435/118](#)

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KWIC	Draw. De
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☐ 23. Document ID: US 20020049217 A1

L4: Entry 23 of 57

File: PGPB

Apr 25, 2002

PGPUB-DOCUMENT-NUMBER: 20020049217

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20020049217 A1

TITLE: Inhibitors of prenyl-protein transferase

PUBLICATION-DATE: April 25, 2002

## INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
deSolms, S. Jane	Collegeville	PA	US	
MacTough, Suzanne C.	Chalfont	PA	US	
Shaw, Anthony W.	Lansdale	PA	US	

US-CL-CURRENT: 514/257; 514/183, 514/397, 514/9, 540/454

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KWIC	Draw. De
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☐ 24. Document ID: US 20020045759 A1

L4: Entry 24 of 57

File: PGPB

Apr 18, 2002

PGPUB-DOCUMENT-NUMBER: 20020045759

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20020045759 A1

TITLE: INHIBITORS OF PRENYL-PROTEIN TRANSFERASE

PUBLICATION-DATE: April 18, 2002

## INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Stump, Craig A.	Pottstown	PA	US	
Williams, Theresa M.	Harleysville	PA	US	

US-CL-CURRENT: 548/153; 548/218, 548/303.1

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KWIC	Draw. De
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☐ 25. Document ID: US 20020037888 A1

L4: Entry 25 of 57

File: PGPB

Mar 28, 2002

PGPUB-DOCUMENT-NUMBER: 20020037888  
PGPUB-FILING-TYPE: new  
DOCUMENT-IDENTIFIER: US 20020037888 A1

TITLE: Inhibitors of prenyl-protein transferase

PUBLICATION-DATE: March 28, 2002

INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
deSolms, S. Jane	Norristown	PA	US	
Graham, Samuel L.	Schwenksville	PA	US	
Shaw, Anthony W.	Lansdale	PA	US	
Ciccarone, Terrence M.	Telford	PA	US	
Stokker, Gerald E.	Gwynedd Valley	PA	US	

US-CL-CURRENT: 514/211.01; 514/211.08, 514/228.8, 514/229.2, 540/544, 540/545,  
544/63, 544/66

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KWIC	Draw. De
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☐ 26. Document ID: US 20020028826 A1

L4: Entry 26 of 57

File: PGPB

Mar 7, 2002

PGPUB-DOCUMENT-NUMBER: 20020028826  
PGPUB-FILING-TYPE: new  
DOCUMENT-IDENTIFIER: US 20020028826 A1

TITLE: HMG-CoA reductase inhibitors and method

PUBLICATION-DATE: March 7, 2002

INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Robl, Jeffrey A.	Newtown	PA	US	
Chen, Bang-Chi	Plainsboro	NJ	US	
Sun, Chong-Qing	East Windsor	NJ	US	

US-CL-CURRENT: 514/290; 546/79, 546/93

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KWIC	Draw. De
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☐ 27. Document ID: US 20020022633 A1

L4: Entry 27 of 57

File: PGPB

Feb 21, 2002

PGPUB-DOCUMENT-NUMBER: 20020022633  
PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20020022633 A1

TITLE: Inhibitors of prenyl-protein transferase

PUBLICATION-DATE: February 21, 2002

## INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Williams, Theresa M.	Harleysville	PA	US	
Stump, Craig A.	Pottstown	PA	US	

US-CL-CURRENT: 514/279; 540/472

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KWIC	Draw. De
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☐ 28. Document ID: US 20020013334 A1

L4: Entry 28 of 57

File: PGPB

Jan 31, 2002

PGPUB-DOCUMENT-NUMBER: 20020013334

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20020013334 A1

TITLE: HMG-CoA reductase inhibitors and method

PUBLICATION-DATE: January 31, 2002

## INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Robl, Jeffrey A.	Newton	PA	US	
Chen, Bang-Chi	Plainsboro	NJ	US	
Sun, Chong-Qing	East Windsor	NJ	US	

US-CL-CURRENT: 514/291; 546/80

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KWIC	Draw. De
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☐ 29. Document ID: US 20020010184 A1

L4: Entry 29 of 57

File: PGPB

Jan 24, 2002

PGPUB-DOCUMENT-NUMBER: 20020010184

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20020010184 A1

TITLE: Inhibitors of prenyl-protein transferase

PUBLICATION-DATE: January 24, 2002

## INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Dinsmore, Christopher J.	Schwenksville	PA	US	
Bergman, Jeffrey M.	Perkasie	PA	US	

US-CL-CURRENT: 514/253.09; 514/254.05, 544/360, 544/370

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KMC	Draw. De
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☐ 30. Document ID: US 6632818 B2

L4: Entry 30 of 57

File: USPT

Oct 14, 2003

US-PAT-NO: 6632818

DOCUMENT-IDENTIFIER: US 6632818 B2

TITLE: Inhibitors of prenyl-protein transferase

DATE-ISSUED: October 14, 2003

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Dinsmore, Christopher J.	Schwenksville	PA		

US-CL-CURRENT: 514/250; 540/456

## ABSTRACT:

The present invention is directed to piperazine-containing macrocyclic compounds which inhibit prenyl-protein transferase, such as farnesyl-protein transferase (FTase), and therefore inhibit the prenylation of the oncogene protein Ras. The invention is further directed to chemotherapeutic compositions containing the compounds of this invention and methods for inhibiting prenyl-protein transferase and the prenylation of the oncogene protein Ras.

15 Claims, 0 Drawing figures

Exemplary Claim Number: 1

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KMC	Draw. De
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☐ 31. Document ID: US 6627636 B2

L4: Entry 31 of 57

File: USPT

Sep 30, 2003

US-PAT-NO: 6627636

DOCUMENT-IDENTIFIER: US 6627636 B2

TITLE: HMG-CoA reductase inhibitors and method

DATE-ISSUED: September 30, 2003



## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Robl; Jeffrey A.	Newtown	PA		

US-CL-CURRENT: 514/291; 514/213.01, 514/292, 540/577, 546/80, 546/81, 546/89,  
546/93

## ABSTRACT:

Compounds of the following structure are HMG CoA reductase inhibitors and thus are active in inhibiting cholesterol biosynthesis, modulating blood serum lipids such as lowering LDL cholesterol and/or increasing HDL cholesterol, and treating hyperlipidemia, hypercholesterolemia, hypertriglyceridemia and atherosclerosis  
##STR1##

and pharmaceutically acceptable salts thereof, wherein X is O, S, SO, SO.sub.2 or NR.sub.7 ; Z is ##STR2##

n is 0 or 1; R.sub.1 and R.sub.2 are the same or different and are independently selected from alkyl, arylalkyl, cycloalkyl, alkenyl, cycloalkenyl, aryl, heteroaryl or cycloheteroalkyl; and R.sub.3 to R.sub.10 are as defined herein.

25 Claims, 0 Drawing figures

Exemplary Claim Number: 1

Full	Title	Citation	Front	Review	Classification	Date	Reference	Abstract	Claims	Keywords	Drawings
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☐ 32. Document ID: US 6620821 B2

L4: Entry 32 of 57

File: USPT

Sep 16, 2003

US-PAT-NO: 6620821

DOCUMENT-IDENTIFIER: US 6620821 B2

TITLE: HMG-CoA reductase inhibitors and method

DATE-ISSUED: September 16, 2003

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Robl; Jeffrey A.	Newtown	PA		

US-CL-CURRENT: 514/290; 546/101, 546/111, 546/93

## ABSTRACT:

Compounds of the following structure are HMG CoA reductase inhibitors and thus are active in inhibiting cholesterol biosynthesis, modulating blood serum lipids, for example, lowering LDL cholesterol and/or increasing HDL cholesterol, and treating hyperlipidemia, dyslipidemia, hormone replacement therapy, hypercholesterolemia, hypertriglyceridemia and atherosclerosis as well as Alzheimer's disease and osteoporosis ##STR1##

and pharmaceutically acceptable salts thereof, ##STR2## n is 0 or 1; x is 0, 1, 2, 3 or 4; y is 0, 1, 2, 3 or 4, provided that at least one of x and y is other than 0; and optionally one or more carbons of (CH.sub.2).sub.x and/or (CH.sub.2).sub.y together with additional carbons form a 3 to 7 membered spirocyclic ring; R.sub.1 and R.sub.2 are the same or different and are independently selected from alkyl, arylalkyl, cycloalkyl, alkenyl, cycloalkenyl, aryl, heteroaryl or cycloheteroalkyl; R.sub.3 is H or lower alkyl; R.sub.4 and R.sub.7 are as defined herein.

18 Claims, 0 Drawing figures  
Exemplary Claim Number: 1

Full	Title	Citation	Front	Review	Classification	Date	Reference	Abstract	Claims	KWC	Draw. De
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☐ 33. Document ID: US 6610722 B2

L4: Entry 33 of 57

File: USPT

Aug 26, 2003

US-PAT-NO: 6610722

DOCUMENT-IDENTIFIER: US 6610722 B2

TITLE: Inhibitors of prenyl-protein transferase

DATE-ISSUED: August 26, 2003

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Stump; Craig A.	Pottstown	PA		
Williams; Theresa M.	Harleysville	PA		

US-CL-CURRENT: 514/397; 540/456, 540/457

ABSTRACT:

The present invention is directed to peptidomimetic macrocyclic compounds that inhibit prenyl-protein transferase and the prenylation of the oncogene protein Ras. The invention is further directed to chemotherapeutic compositions containing the compounds of this invention and methods for inhibiting prenyl-protein transferase and the prenylation of the oncogene protein Ras.

16 Claims, 0 Drawing figures  
Exemplary Claim Number: 1

Full	Title	Citation	Front	Review	Classification	Date	Reference	Abstract	Claims	KWC	Draw. De
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☐ 34. Document ID: US 6583290 B1

L4: Entry 34 of 57

File: USPT

Jun 24, 2003

US-PAT-NO: 6583290

DOCUMENT-IDENTIFIER: US 6583290 B1

TITLE: 14-methyl epothilone derivatives

DATE-ISSUED: June 24, 2003

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Julien; Bryan	Oakland	CA		
Katz; Leonard	Hayward	CA		
Khosla; Chaitan	Palo Alto	CA		
Tang; Li	Foster City	CA		
Ziermann; Rainer	San Mateo	CA		

US-CL-CURRENT: 548/203; 181/205, 546/268.1

## ABSTRACT:

Compounds of the invention include 14-methyl epothilone derivatives. More generally, preferred compounds of the invention are those that can be produced by altering the epothilone PKS genes as described herein and optionally by action of epothilone modification enzymes and/or by chemically modifying the resulting epothilones produces when those genes are expressed.

2 Claims, 9 Drawing figures

Exemplary Claim Number: 1

Number of Drawing Sheets: 8

Full	Title	Citation	Front	Review	Classification	Date	Reference	Abstract	Claims	MM	Draw. Data
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☐ 35. Document ID: US 6566385 B2

L4: Entry 35 of 57

File: USPT

May 20, 2003

US-PAT-NO: 6566385

DOCUMENT-IDENTIFIER: US 6566385 B2

TITLE: Inhibitors of prenyl-protein transferase

DATE-ISSUED: May 20, 2003

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
deSolms; S. Jane	Collegeville	PA		
Stokker; Gerald E.	Gwynedd Valley	PA		
Shaw; Anthony W.	Lansdale	PA		

US-CL-CURRENT: 514/397; 540/456, 540/468, 548/311.7

## ABSTRACT:

The present invention is directed to peptidomimetic macrocyclic compounds which inhibit prenyl-protein transferase and the prenylation of the oncogene protein Ras.

The invention is further directed to chemotherapeutic compositions containing the compounds of this invention and methods for inhibiting prenyl-protein transferase and the prenylation of the oncogene protein Ras.

22 Claims, 0 Drawing figures  
Exemplary Claim Number: 1

Full	Title	Citation	Front	Review	Classification	Date	Reference	Abstract	Claims	Keywords	Drawings
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☐ 36. Document ID: US 6562823 B1

L4: Entry 36 of 57

File: USPT

May 13, 2003

US-PAT-NO: 6562823

DOCUMENT-IDENTIFIER: US 6562823 B1

TITLE: Inhibitors of prenyl-protein transferase

DATE-ISSUED: May 13, 2003

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Dinsmore; Christopher J.	Schwenksville	PA		
Bergman; Jeffrey M.	Perkasie	PA		
Graham; Samuel L.	Schwenksville	PA		
Nguyen; Diem N.	North Wales	PA		
Stokker; Gerald E.	Gwynedd Valley	PA		
Williams; Theresa M.	Harleysville	PA		
Zartman; C. Blair	Hatfield	PA		

US-CL-CURRENT: 514/250; 540/469

ABSTRACT:

The present invention is directed to peptidomimetic piperazine-containing macrocyclic compounds which inhibit prenyl-protein transferase and the prenylation of the oncogene protein Ras. The invention is further directed to chemotherapeutic compositions containing the compounds of this invention and methods for inhibiting prenyl-protein transferase and the prenylation of the oncogene protein Ras.

17 Claims, 0 Drawing figures  
Exemplary Claim Number: 1

Full	Title	Citation	Front	Review	Classification	Date	Reference	Abstract	Claims	Keywords	Drawings
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☐ 37. Document ID: US 6534506 B2

L4: Entry 37 of 57

File: USPT

Mar 18, 2003

US-PAT-NO: 6534506

DOCUMENT-IDENTIFIER: US 6534506 B2

TITLE: Inhibitors of prenyl-protein transferase

DATE-ISSUED: March 18, 2003

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Nguyen; Diem N.	North Wales	PA		
Stump; Craig A.	Pottstown	PA		
Williams; Theresa M.	Harleysville	PA		

US-CL-CURRENT: 514/250; 540/456, 540/460, 540/469, 540/472, 544/229, 544/231,  
544/362, 544/368, 544/370, 548/110, 548/154, 548/218, 548/322.5

## ABSTRACT:

The present invention is directed to peptidomimetic macrocyclic compounds which inhibit prenyl-protein transferase and the prenylation of the oncogene protein Ras. The invention is further directed to chemotherapeutic compositions containing the compounds of this invention and methods for inhibiting prenyl-protein transferase and the prenylation of the oncogene protein Ras.

17 Claims, 0 Drawing figures

Exemplary Claim Number: 1

Full	Title	Citation	Front	Review	Classification	Date	Reference			Claims	KWIC	Draw. De
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☐ 38. Document ID: US 6525074 B2

L4: Entry 38 of 57

File: USPT

Feb 25, 2003

US-PAT-NO: 6525074

DOCUMENT-IDENTIFIER: US 6525074 B2

TITLE: Inhibitors of prenyl-protein transferase

DATE-ISSUED: February 25, 2003

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
deSolms; S. Jane	Collegeville	PA		
MacTough; Suzanne C.	Chalfont	PA		
Shaw; Anthony W.	Lansdale	PA		

US-CL-CURRENT: 514/338; 514/397, 540/456, 540/472

## ABSTRACT:

The present invention is directed to peptidomimetic macrocyclic compounds which

inhibit prenyl-protein transferase and the prenylation of the oncogene protein Ras. The invention is further directed to chemotherapeutic compositions containing the compounds of this invention and methods for inhibiting prenyl-protein transferase and the prenylation of the oncogene protein Ras.

28 Claims, 0 Drawing figures

Exemplary Claim Number: 1

Full	Title	Citation	Front	Review	Classification	Date	Reference	Abstract	Pub. No.	Claims	KWC	Draw. De
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☐ 39. Document ID: US 6441017 B1

L4: Entry 39 of 57

File: USPT

Aug 27, 2002

US-PAT-NO: 6441017

DOCUMENT-IDENTIFIER: US 6441017 B1

TITLE: Inhibitors of prenyl-protein transferase

DATE-ISSUED: August 27, 2002

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Bell; Ian M.	Harleysville	PA		
Beshore; Douglas C.	Lansdale	PA		
Gallicchio; Steven N.	Ambler	PA		
Zartman; C. Blair	Hatfield	PA		

US-CL-CURRENT: 514/393; 540/456, 540/472

ABSTRACT:

The present invention is directed to macrocyclic compounds which inhibit prenyl-protein transferase (FTase) and the prenylation of the oncogene protein Ras. The invention is further directed to chemotherapeutic compositions containing the compounds of this invention and methods for inhibiting prenyl-protein transferase and the prenylation of the oncogene protein Ras.

26 Claims, 0 Drawing figures

Exemplary Claim Number: 1

Full	Title	Citation	Front	Review	Classification	Date	Reference	Abstract	Pub. No.	Claims	KWC	Draw. De
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☐ 40. Document ID: US 6413964 B1

L4: Entry 40 of 57

File: USPT

Jul 2, 2002

US-PAT-NO: 6413964

DOCUMENT-IDENTIFIER: US 6413964 B1

TITLE: Inhibitors of prenyl-protein transferase

DATE-ISSUED: July 2, 2002

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
deSolms; S. Jane	Rahway	NJ	07065-0907	
MacTough; Suzanne C.	Rahway	NJ	07065-0907	
Shaw; Anthony W.	Rahway	NJ	07065-0907	

US-CL-CURRENT: [514/250](#); [540/456](#), [540/458](#), [540/460](#), [540/469](#), [540/472](#)

## ABSTRACT:

The present invention is directed to peptidomimetic macrocyclic compounds of the formula A: ##STR1##

wherein W is a heterocycle, V is a heterocycle or aryl moiety and Z<sup>sup.1</sup> is a suitably substituted aryl or heterocycle moiety. The instant compounds inhibit prenyl-protein transferase and the prenylation of the oncogene protein Ras. The invention is further directed to chemotherapeutic compositions containing the compounds of this invention and methods for inhibiting prenyl-protein transferase and the prenylation of the oncogene protein Ras.

18 Claims, 0 Drawing figures

Exemplary Claim Number: 1

Full	Title	Citation	Front	Review	Classification	Date	Reference	Abstract	Full Text	Claims	RWC	Draw. Data
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☐ 41. Document ID: US 6410534 B1

**Using default format because multiple data bases are involved.**

L4: Entry 41 of 57

File: USPT

Jun 25, 2002

US-PAT-NO: 6410534

DOCUMENT-IDENTIFIER: US 6410534 B1

TITLE: Inhibitors of prenyl-protein transferase

DATE-ISSUED: June 25, 2002

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Dinsmore; Christopher J.	Schwenksville	PA		
Bell; Ian M.	Harleyville	PA		
Beshore; Douglas C.	Lansdale	PA		
Williams; Theresa M.	Harleysville	PA		

US-CL-CURRENT: 514/249, 514/250, 540/457, 540/458, 540/459, 540/461, 540/468,  
540/469, 540/471, 540/472, 540/476, 540/477

Full	Title	Citation	Front	Review	Classification	Date	Reference	Abstract	Abstract	Claims	Drawings	Drawings
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☐ 42. Document ID: US 6410301 B1

L4: Entry 42 of 57

File: USPT

Jun 25, 2002

US-PAT-NO: 6410301

DOCUMENT-IDENTIFIER: US 6410301 B1

TITLE: Myxococcus host cells for the production of epothilones

DATE-ISSUED: June 25, 2002

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Julien; Bryan	Oakland	CA		
Katz; Leonard	Hayward	CA		
Khosla; Chaitan	Palo Alto	CA		



US-CL-CURRENT: 435/252.3

## ABSTRACT:

Recombinant Myxococcus host cell containing recombinant expression vectors containing epothilone polyketide synthase genes can produce epothilones C and D.

6 Claims, 4 Drawing figures

Exemplary Claim Number: 1

Number of Drawing Sheets: 4

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	FIGS	Draw. De
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☐ 43. Document ID: US 6387903 B1

L4: Entry 43 of 57

File: USPT

May 14, 2002

US-PAT-NO: 6387903

DOCUMENT-IDENTIFIER: US 6387903 B1

TITLE: Inhibitors of prenyl-protein transferase

DATE-ISSUED: May 14, 2002

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Dinsmore; Christopher J	Schwenksville	PA		
Hutchinson; John H.	Philadelphia	PA		
Williams; Theresa M.	Harleysville	PA		

US-CL-CURRENT: 514/254.05; 544/370

## ABSTRACT:

The present invention is directed to compounds which inhibit prenyl-protein transferases, farnesyl-protein transferase and geranylgeranyl-protein transferase type I, and the prenylation of the oncogene protein Ras. The invention is further directed to chemotherapeutic compositions containing the compounds of this invention and methods for inhibiting farnesyl-protein transferase and geranylgeranyl-protein transferase type I and the prenylation of the oncogene protein RAS.

12 Claims, 0 Drawing figures

Exemplary Claim Number: 1

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	FIGS	Draw. De
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☐ 44. Document ID: US 6380228 B1

L4: Entry 44 of 57

File: USPT

Apr 30, 2002

US-PAT-NO: 6380228

DOCUMENT-IDENTIFIER: US 6380228 B1

TITLE: Inhibitors of prenyl-protein transferase

DATE-ISSUED: April 30, 2002

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Stump; Craig A.	Pottstown	PA		
Williams; Theresa M.	Harleysville	PA		

US-CL-CURRENT: 514/368; 514/375, 514/393, 548/154, 548/218, 548/302.7

## ABSTRACT:

The present invention is directed to peptidomimetic compounds which inhibit prenyl-protein transferase and the prenylation of the oncogene protein Ras. The invention is further directed to chemotherapeutic compositions containing the compounds of this invention and methods for inhibiting prenyl-protein transferase and the prenylation of the oncogene protein Ras.

22 Claims, 0 Drawing figures

Exemplary Claim Number: 1

Full	Title	Citation	Front	Review	Classification	Date	Reference			Claims	KWC	Draw. D
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☐ 45. Document ID: US 6376496 B1

L4: Entry 45 of 57

File: USPT

Apr 23, 2002

US-PAT-NO: 6376496

DOCUMENT-IDENTIFIER: US 6376496 B1

TITLE: Inhibitors of prenyl-protein transferase

DATE-ISSUED: April 23, 2002

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Hartman; George D.	Lansdale	PA		
Lumma, Jr.; William C.	Pennsburg	PA		
Sisko; John T.	Lansdale	PA		
Smith; Anthony M.	Green Lane	PA		
Tucker; Thomas J.	North Wales	PA		
Bergman; Jeffrey M.	Perkasie	PA		

US-CL-CURRENT: 514/254.05; 544/370

## ABSTRACT:

The present invention comprises piperazine-containing compounds which inhibit prenyl-protein transferases, including farnesyl-protein transferase and geranylgeranyl-protein transferase type I. Such therapeutic compounds are useful in the treatment of cancer.

33 Claims, 0 Drawing figures

Exemplary Claim Number: 1

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	RMIC	Draw. De
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☐ 46. Document ID: US 6358985 B1

L4: Entry 46 of 57

File: USPT

Mar 19, 2002

US-PAT-NO: 6358985

DOCUMENT-IDENTIFIER: US 6358985 B1

**\*\* See image for Certificate of Correction \*\***

TITLE: Inhibitors of prenyl-protein transferase

DATE-ISSUED: March 19, 2002

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Anthony; Neville J.	Hatfield	PA		
Bell; Ian M.	Harleysville	PA		
Beshore; Douglas C.	Lansdale	PA		
Ciccarone; Terrence M.	Telford	PA		
de Solms; S. Jane	Norristown	PA		
Dinsmore; Christopher J.	Schwenksville	PA		
Stokker; Gerald E.	Gwynedd Valley	PA		

US-CL-CURRENT: 514/393; 540/455, 540/456, 540/468

ABSTRACT:

The present invention is directed to peptidomimetic macrocyclic compounds which inhibit prenyl-protein transferase and the prenylation of the oncogene protein Ras. The invention is further directed to chemotherapeutic compositions containing the compounds of this invention and methods for inhibiting prenyl-protein transferase and the prenylation of the oncogene protein Ras.

26 Claims, 0 Drawing figures

Exemplary Claim Number: 1

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	RMIC	Draw. De
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☐ 47. Document ID: US 6358956 B1

L4: Entry 47 of 57

File: USPT

Mar 19, 2002

US-PAT-NO: 6358956  
DOCUMENT-IDENTIFIER: US 6358956 B1

TITLE: Inhibitors of prenyl-protein transferase

DATE-ISSUED: March 19, 2002

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Hartman; George D.	Lansdale	PA		
Lumma, Jr.; William C.	Pennsburg	PA		
Sisko; John T.	Lansdale	PA		
Smith; Anthony M.	Green Lane	PA		
Tucker; Thomas J.	North Wales	PA		
Stokker; Gerald E.	Gwynedd Valley	PA		

US-CL-CURRENT: 514/252.13; 544/358, 544/366, 544/379, 544/389, 544/391, 548/311.1,  
548/314.7

ABSTRACT:

The present invention comprises piperazine-containing compounds which inhibit prenyl-protein transferases, including-farnesyl-protein transferase and geranylgeranyl-protein transferase type I. Such therapeutic compounds are useful in the treatment of cancer.

22 Claims, 0 Drawing figures  
Exemplary Claim Number: 1

Full	Title	Citation	Front	Review	Classification	Date	Reference			Claims	KWIC	Draw. De
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☐ 48. Document ID: US 6355643 B1

L4: Entry 48 of 57

File: USPT

Mar 12, 2002

US-PAT-NO: 6355643  
DOCUMENT-IDENTIFIER: US 6355643 B1

**\*\* See image for Certificate of Correction \*\***

TITLE: Inhibitors of prenyl-protein transferase

DATE-ISSUED: March 12, 2002

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Lumma; William C.	Pennsburg	PA		
Sisko; John T.	Lansdale	PA		
Smith; Anthony M.	Green Lane	PA		
Tucker; Thomas J.	North Wales	PA		
Dinsmore; Christopher J.	Schwenksville	PA		

Bergman; Jeffrey M.

Perkasie

PA

US-CL-CURRENT: 514/254.05; 514/218, 514/253.06, 514/253.09, 514/253.1, 540/575,  
544/231, 544/363, 544/364, 544/367, 544/369, 544/370

## ABSTRACT:

The present invention comprises unsubstituted and substituted piperazine-containing compounds having alkanoyl, alkylsulfonyl, alkylamido or alkoxy carbonyl substituents, and having the formula A, ##STR1##

which inhibit prenyl-protein transferases. In particular, the invention relates to prenyl-protein transferase inhibitors which are efficacious in vivo as inhibitors of geranylgeranyl-protein transferase type I (GGTase-I) and that inhibit the cellular processing of both the H-Ras protein and the K4B-Ras protein. Such therapeutic compounds are useful in the treatment of cancer.

19 Claims, 0 Drawing figures  
Exemplary Claim Number: 1,2

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWIC	Draw. De
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☐ 49. Document ID: US 6350755 B1

L4: Entry 49 of 57

File: USPT

Feb 26, 2002

US-PAT-NO: 6350755

DOCUMENT-IDENTIFIER: US 6350755 B1

TITLE: Inhibitors of prenyl-protein transferase

DATE-ISSUED: February 26, 2002

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
deSolms; S. Jane	Collegeville	PA		
Shaw; Anthony W.	Lansdale	PA		

US-CL-CURRENT: 514/281; 540/456

## ABSTRACT:

The present invention is directed to peptidomimetic macrocyclic compounds which inhibit prenyl-protein transferase and the prenylation of the oncogene protein Ras. The invention is further directed to chemotherapeutic compositions containing the compounds of this invention and methods for inhibiting prenyl-protein transferase and the prenylation of the oncogene protein Ras.

18 Claims, 0 Drawing figures  
Exemplary Claim Number: 1

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWIC	Draw De
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☐ 50. Document ID: US 6335343 B1

L4: Entry 50 of 57

File: USPT

Jan 1, 2002

US-PAT-NO: 6335343

DOCUMENT-IDENTIFIER: US 6335343 B1

TITLE: Inhibitors of prenyl-protein transferase

DATE-ISSUED: January 1, 2002

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Lumma, Jr.; William C.	Pennsburg	PA		
Smith; Anthony M.	Green Lane	PA		
Sisko; John T.	Lansdale	PA		

US-CL-CURRENT: 514/254.05; 514/218, 514/253.06, 514/253.07, 514/253.09, 514/254.02,  
514/254.04, 540/575, 544/231, 544/363, 544/364, 544/367, 544/369, 544/370

## ABSTRACT:

The present invention comprises piperazine/piperazinone-containing compounds having multicyclic ring system substituents on one of the piperazine/piperazinone nitrogens, which inhibit prenyl-protein transferases, including farnesyl-protein transferase and geranylgeranyl-protein transferase type I. Such therapeutic compounds are useful in the treatment of cancer.

18 Claims, 0 Drawing figures

Exemplary Claim Number: 1,2

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWIC	Draw De
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☐ 51. Document ID: US 6333335 B1

L4: Entry 51 of 57

File: USPT

Dec 25, 2001

US-PAT-NO: 6333335

DOCUMENT-IDENTIFIER: US 6333335 B1

TITLE: Phenyl-protein transferase inhibitors

DATE-ISSUED: December 25, 2001

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Dinsmore; Christopher J.	Schwenksville	PA		
Graham; Samuel L.	Schwenksville	PA		

Williams; Theresa M.                      Harleysville                      PA

US-CL-CURRENT: 514/300; 546/121

ABSTRACT:

The present invention is directed to tetrahydro-imidazopyridinyl compounds which inhibit prenyl-protein transferase and the prenylation of the oncogene protein Ras. The invention is further directed to chemotherapeutic compositions containing the compounds of this invention and methods for inhibiting prenyl-protein transferase and the prenylation of the oncogene protein Ras.

6 Claims, 0 Drawing figures  
Exemplary Claim Number: 1

Full	Title	Citation	Front	Review	Classification	Date	Reference	Abstract	Claims	KWIC	Draw De
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☐ 52. Document ID: US 6329376 B1

L4: Entry 52 of 57

File: USPT

Dec 11, 2001

US-PAT-NO: 6329376

DOCUMENT-IDENTIFIER: US 6329376 B1

TITLE: Inhibitors of prenyl-protein transferase

DATE-ISSUED: December 11, 2001

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Bergman; Jeffrey M.	Perkasie	PA		

US-CL-CURRENT: 514/250; 540/469

ABSTRACT:

The present invention is directed to peptidomimetic piperazine-containing macrocyclic compounds which inhibit a prenyl-protein transferase (FTase) and the prenylation of the oncogene protein Ras. The invention is further directed to chemotherapeutic compositions containing the compounds of this invention and methods for inhibiting prenyl-protein transferase and the prenylation of the oncogene protein Ras.

14 Claims, 0 Drawing figures  
Exemplary Claim Number: 1

Full	Title	Citation	Front	Review	Classification	Date	Reference	Abstract	Claims	KWIC	Draw De
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☐ 53. Document ID: US 6316436 B1

L4: Entry 53 of 57

File: USPT

Nov 13, 2001

US-PAT-NO: 6316436

DOCUMENT-IDENTIFIER: US 6316436 B1

TITLE: Inhibitors of prenyl-protein transferase

DATE-ISSUED: November 13, 2001

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
deSolms; S. Jane	Norristown	PA		
Shaw; Anthony W.	Lansdale	PA		

US-CL-CURRENT: 514/211.1; 540/469

## ABSTRACT:

The present invention is directed to compounds which inhibit prenyl-protein transferase and particularly, the farnesylation of the oncogene protein Ras. The invention is further directed to chemotherapeutic compositions containing the compounds of this invention and methods for inhibiting farnesyl-protein transferase and the farnesylation of the oncogene protein Ras.

17 Claims, 0 Drawing figures

Exemplary Claim Number: 1

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWIC	Draw. De
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☐ 54. Document ID: US 6303342 B1

L4: Entry 54 of 57

File: USPT

Oct 16, 2001

US-PAT-NO: 6303342

DOCUMENT-IDENTIFIER: US 6303342 B1

TITLE: Recombinant methods and materials for producing epothilones C and D

DATE-ISSUED: October 16, 2001

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Julien; Bryan	Oakland	CA		
Katz; Leonard	Hayward	CA		
Khosla; Chaitan	Palo Alto	CA		
Tang; Li	Foster City	CA		

US-CL-CURRENT: 435/76

## ABSTRACT:



Recombinant nucleic acids that encode all or a portion of the epothilone polyketide synthase (PKS) are used to express recombinant PKS genes in host cells for the production of epothilones, epothilone derivatives, and polyketides that are useful as cancer chemotherapeutics, fungicides, and immunosuppressants.

29 Claims, 9 Drawing figures  
Exemplary Claim Number: 1  
Number of Drawing Sheets: 8

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KMC	Draw. De
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☐ 55. Document ID: US 6297239 B1

L4: Entry 55 of 57

File: USPT

Oct 2, 2001

US-PAT-NO: 6297239

DOCUMENT-IDENTIFIER: US 6297239 B1

**\*\* See image for Certificate of Correction \*\***

TITLE: Inhibitors of prenyl-protein transferase

DATE-ISSUED: October 2, 2001

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
deSolms; S. Jane	Norristown	PA		
Hutchinson; John H.	Philadelphia	PA		
Shaw; Anthony W.	Lansdale	PA		
Graham; Samuel L.	Schwenksville	PA		
Ciccarone; Terrence M.	Telford	PA		

US-CL-CURRENT: 514/235.8; 514/399, 544/122, 548/335.1, 548/336.1, 548/346.1

ABSTRACT:

The present invention is directed to compounds which inhibit a prenyl-protein transferase (FTase) and the farnesylation of the oncogene protein Ras. The invention is further directed to chemotherapeutic compositions containing the compounds of this invention and methods for inhibiting a prenyl-protein transferase and the prenylation of the oncogene protein Ras.

31 Claims, 0 Drawing figures  
Exemplary Claim Number: 1

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KMC	Draw. De
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☐ 56. Document ID: US 6284755 B1

L4: Entry 56 of 57

File: USPT

Sep 4, 2001

US-PAT-NO: 6284755

DOCUMENT-IDENTIFIER: US 6284755 B1

**\*\* See image for Certificate of Correction \*\***

TITLE: Inhibitors of prenyl-protein transferase

DATE-ISSUED: September 4, 2001

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
deSolms; S. Jane	Norristown	PA		
Graham; Samuel L.	Schwenksville	PA		
Shaw; Anthony W.	Lansdale	PA		
Ciccarone; Terrence M.	Telford	PA		
Stokker; Gerald E.	Gwynedd Valley	PA		

US-CL-CURRENT: 514/212.03; 514/212.08, 540/524, 540/531

## ABSTRACT:

The present invention is directed to azepan-2-one compounds which inhibit prenyl-protein transferase, particularly farnesyl-protein transferase (Ftase), and the prenylation of the oncogene protein Ras. The invention is further directed to chemotherapeutic compositions containing the compounds of this invention and methods for inhibiting prenyl-protein transferase and the prenylation of the oncogene protein Ras.

35 Claims, 0 Drawing figures

Exemplary Claim Number: 1

Full	Title	Citation	Front	Review	Classification	Date	Reference	Abstract	Claims	KWIC	Draw De
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☐ 57. Document ID: US 6127390 A

L4: Entry 57 of 57

File: USPT

Oct 3, 2000

US-PAT-NO: 6127390

DOCUMENT-IDENTIFIER: US 6127390 A

TITLE: Inhibitors of prenyl-protein transferase

DATE-ISSUED: October 3, 2000

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
deSolms; S. Jane	Norristown	PA		
Lumma, Jr.; William C.	Pennsburg	PA		
Shaw; Anthony W.	Lansdale	PA		
Sisko; John T.	Lansdale	PA		
Tucker; Thomas J.	North Wales	PA		

US-CL-CURRENT: [514/341](#); [546/274.1](#), [546/274.4](#), [546/275.1](#)

ABSTRACT:

The present invention is directed to compounds which inhibit prenyl-protein transferase (FTase) and the prenylation of the oncogene protein Ras. The invention is further directed to chemotherapeutic compositions containing the compounds of this invention and methods for inhibiting prenyl-protein transferase and the prenylation of the oncogene protein Ras.

18 Claims, 0 Drawing figures

Exemplary Claim Number: 1

Full	Title	Citation	Front	Review	Classification	Date	Reference	Abstract	Attached	Claims	KWOC	Draw. Data
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Terms	Documents
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**Search Results - Record(s) 1 through 20 of 40 returned.**☐ 1. Document ID: US 20040082026 A1**Using default format because multiple data bases are involved.**

L7: Entry 1 of 40

File: PGPB

Apr 29, 2004

PGPUB-DOCUMENT-NUMBER: 20040082026

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20040082026 A1

TITLE: Interferon alpha: remodeling and glycoconjugation of interferon alpha

PUBLICATION-DATE: April 29, 2004

## INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
DeFrees, Shawn	North Wales	PA	US	
Zopf, David	Wayne	PA	US	
Bayer, Robert	San Diego	CA	US	
Bowe, Caryn	Doylestown	PA	US	
Hakes, David	Willow Grove	PA	US	
Chen, Xi	Lansdale	PA	US	

US-CL-CURRENT: 435/68.1; 530/351

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	RWC	Draw D
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☐ 2. Document ID: US 20040077836 A1

L7: Entry 2 of 40

File: PGPB

Apr 22, 2004

PGPUB-DOCUMENT-NUMBER: 20040077836

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20040077836 A1

TITLE: Granulocyte colony stimulating factor: remodeling and glycoconjugation of G-CSF

PUBLICATION-DATE: April 22, 2004

## INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
------	------	-------	---------	---------

DeFrees, Shawn	North Wales	PA	US
Zopf, David	Wayne	PA	US
Bayer, Robert	San Diego	CA	US
Bowe, Caryn	Doylestown	PA	US
Hakes, David	Willow Grove	PA	US
Chen, Xi	Lansdale	PA	US

US-CL-CURRENT: 530/351; 435/68.1

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KMC	Draw D
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☐ 3. Document ID: US 20040063911 A1

L7: Entry 3 of 40

File: PGPB

Apr 1, 2004

PGPUB-DOCUMENT-NUMBER: 20040063911

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20040063911 A1

TITLE: Protein remodeling methods and proteins/peptides produced by the methods

PUBLICATION-DATE: April 1, 2004

## INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
DeFrees, Shawn	North Wales	PA	US	
Zopf, David	Wayne	PA	US	
Bayer, Robert	San Diego	CA	US	
Hakes, David	Willow Grove	PA	US	
Chen, Xi	Lansdale	PA	US	

US-CL-CURRENT: 530/351; 435/68.1, 530/395

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KMC	Draw D
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☐ 4. Document ID: US 20040053995 A1

L7: Entry 4 of 40

File: PGPB

Mar 18, 2004

PGPUB-DOCUMENT-NUMBER: 20040053995

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20040053995 A1

TITLE: Synthesis of epothilones, intermediates thereto and analogues thereof

PUBLICATION-DATE: March 18, 2004

## INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
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Danishefsky, Samuel J.	Englewood	NJ	US
Rivkin, Alexey	New York	NY	US
Yoshimura, Fumikiko	New York	NY	US
Chou, Ting-Chao	Paramus	NJ	US
Gabarda, Ana E.	New York	NY	US

US-CL-CURRENT: [514/465](#); [514/474](#), [540/451](#), [548/203](#), [548/235](#)

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KWIC	Draw D
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☐ 5. Document ID: US 20040053910 A1

L7: Entry 5 of 40

File: PGPB

Mar 18, 2004

PGPUB-DOCUMENT-NUMBER: 20040053910

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20040053910 A1

TITLE: Synthesis of epothilones, intermediates thereto and analogues thereof

PUBLICATION-DATE: March 18, 2004

## INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Danishefsky, Samuel J.	Englewood	NJ	US	
Rivkin, Alexey	New York	NY	US	
Fumikiko, Yoshimura	New York	NY	US	
Chou, Ting-Chao	Paramus	NJ	US	
Gabarda, Ana E.	New York	NY	US	
Dong, Huajin	New York	NY	US	

US-CL-CURRENT: [514/183](#); [514/23](#), [514/365](#), [514/431](#), [514/450](#), [536/17.4](#), [540/451](#), [548/181](#)

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KWIC	Draw D
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☐ 6. Document ID: US 20040043446 A1

L7: Entry 6 of 40

File: PGPB

Mar 4, 2004

PGPUB-DOCUMENT-NUMBER: 20040043446

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20040043446 A1

TITLE: Alpha galactosidase a: remodeling and glycoconjugation of alpha galactosidase A

PUBLICATION-DATE: March 4, 2004

## INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
DeFrees, Shawn	North Wales	PA	US	
Zopf, David	Wayne	PA	US	
Bayer, Robert	San Diego	CA	US	
Bowe, Caryn	Doylestown	PA	US	
Hakes, David	Willow Grove	PA	US	
Chen, Xi	Lansdale	PA	US	

US-CL-CURRENT: [435/68.1](#); [435/193](#), [435/208](#)

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KWOC	Draw. D
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☐ 7. Document ID: US 20040030147 A1

L7: Entry 7 of 40

File: PGPB

Feb 12, 2004

PGPUB-DOCUMENT-NUMBER: 20040030147  
PGPUB-FILING-TYPE: new  
DOCUMENT-IDENTIFIER: US 20040030147 A1

TITLE: Method for synthesizing epothilones and epothilone analogs

PUBLICATION-DATE: February 12, 2004

## INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
White, James David	Philomath	OR	US	
Sundermann, Kurt Frederick	Burlingame	CA	US	
Carter, Rich Garrett	Corvallis	OR	US	

US-CL-CURRENT: [548/181](#); [548/234](#)

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KWOC	Draw. D
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☐ 8. Document ID: US 20040014183 A1

L7: Entry 8 of 40

File: PGPB

Jan 22, 2004

PGPUB-DOCUMENT-NUMBER: 20040014183  
PGPUB-FILING-TYPE: new  
DOCUMENT-IDENTIFIER: US 20040014183 A1

TITLE: Secondary metabolite congener distribution modulation

PUBLICATION-DATE: January 22, 2004

## INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Licari, Peter J.	Fremont	CA	US	

Julien, Bryan	Oakland	CA	US
Frykman, Scott	Hayward	CA	US
Tsuruta, Hiroko	Emeryville	CA	US

US-CL-CURRENT: 435/118; 435/252.3

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KMC	Draw. De
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☐ 9. Document ID: US 20030219877 A1

L7: Entry 9 of 40

File: PGPB

Nov 27, 2003

PGPUB-DOCUMENT-NUMBER: 20030219877  
PGPUB-FILING-TYPE: new  
DOCUMENT-IDENTIFIER: US 20030219877 A1

TITLE: Novel epothilone compounds and methods for making the same

PUBLICATION-DATE: November 27, 2003

## INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Tang, Li	Foster City	CA	US	
Metcalf, Brian	Moraga	CA	US	
Katz, Leonard	Oakland	CA	US	
Ashley, Gary	Alameda	CA	US	

US-CL-CURRENT: 435/71.1; 435/170

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KMC	Draw. De
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☐ 10. Document ID: US 20030208080 A1

L7: Entry 10 of 40

File: PGPB

Nov 6, 2003

PGPUB-DOCUMENT-NUMBER: 20030208080  
PGPUB-FILING-TYPE: new  
DOCUMENT-IDENTIFIER: US 20030208080 A1

TITLE: Synthesis of epothilones, intermediates thereto and analogues thereof

PUBLICATION-DATE: November 6, 2003

## INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Danishefsky, Samuel J.	Englewood	NJ	US	
Bertinato, Peter	Old Lyme	CT	US	
Su, Dai-Shi	Ambler	PA	US	
Meng, DongFang	New York	NY	US	



Chou, Ting-Chao	Paramus	NJ	US
Kamenecka, Ted	New Brunswick	NJ	US
Sorensen, Erik J.	San Diego	CA	US
Balog, Aaron	Scotch Plains	NJ	US
Savin, Kenneth A.	Indianapolis	IN	US
Kuduk, Scott	Harleysville	PA	US
Harris, Christina	New York	NY	US
Zhang, Xiu-Guo	New York	NY	US
Bertino, Joseph R.	Branford	CT	US

US-CL-CURRENT: [546/281.7](#); [548/204](#), [548/237](#), [548/311.4](#), [548/465](#), [549/266](#)

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KWIC	Draw. De
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☐ 11. Document ID: US 20030187273 A1

L7: Entry 11 of 40

File: PGPB

Oct 2, 2003

PGPUB-DOCUMENT-NUMBER: 20030187273

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20030187273 A1

TITLE: Method for synthesizing epothilones and epothilone analogs

PUBLICATION-DATE: October 2, 2003

## INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
White, James David	Philomath	OR	US	
Carter, Rich Garrett	Oxford	MS	US	
Sundermann, Kurt Frederick	Corvallis	OR	US	

US-CL-CURRENT: [548/204](#); [548/181](#)

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KWIC	Draw. De
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☐ 12. Document ID: US 20030176473 A1

L7: Entry 12 of 40

File: PGPB

Sep 18, 2003

PGPUB-DOCUMENT-NUMBER: 20030176473

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20030176473 A1

TITLE: Derivatives of epothilone B and D and synthesis thereof

PUBLICATION-DATE: September 18, 2003

## INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Taylor, Richard E.	South Bend	IN	US	
Chen, Yue	Mishawaka	IN	US	

US-CL-CURRENT: [514/365](#); [548/181](#), [548/204](#)

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	RMC	Draw. De
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☐ 13. Document ID: US 20030176368 A1

L7: Entry 13 of 40

File: PGPB

Sep 18, 2003

PGPUB-DOCUMENT-NUMBER: 20030176368  
PGPUB-FILING-TYPE: new  
DOCUMENT-IDENTIFIER: US 20030176368 A1

TITLE: Synthesis of epothilones, intermediates thereto and analogues thereof

PUBLICATION-DATE: September 18, 2003

## INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Danishefsky, Samuel J.	Englewood	NJ	US	
Biswas, Kaustav	Thouand Oaks	NY	US	
Chappell, Mark	Noblesville	IN	US	
Lin, Hong	New York	NY	US	
Njardarson, Jon T.	New York	NY	US	
Lee, Chul Bom	Princeton	NJ	US	
Rivkin, Alexy	New York	NY	US	
Chou, Ting-Chao	Paramus	NJ	US	

US-CL-CURRENT: [514/28](#); [514/183](#), [514/365](#), [514/431](#), [514/450](#), [536/7.4](#), [540/451](#),  
[548/203](#), [549/266](#), [549/9](#)

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	RMC	Draw. De
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☐ 14. Document ID: US 20030105330 A1

L7: Entry 14 of 40

File: PGPB

Jun 5, 2003

PGPUB-DOCUMENT-NUMBER: 20030105330  
PGPUB-FILING-TYPE: new  
DOCUMENT-IDENTIFIER: US 20030105330 A1

TITLE: Synthesis of epothilones, intermediates thereto and analogues thereof

PUBLICATION-DATE: June 5, 2003

## INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
------	------	-------	---------	---------

Danishefsky, Samuel J.	Englewood	NJ	US
Bertinato, Peter	Old Lyme	CT	US
Su, Dai-Shi	New York	NY	US
Meng, Dongfang	New York	NY	US
Chou, Ting-Chao	Paramus	NJ	US
Kamenecka, Ted	New York	NY	US
Sorensen, Erik J.	San Diego	CA	US
Balog, Aaron	New York	NY	US
Savin, Kenneth A.	New York	NY	US
Kuduk, Scott	Harleysville	PA	US
Harris, Christina	New York	NY	US
Zhang, Xiu-Guo	New York	NY	US
Bertino, Joseph R.	Branford	CT	US

US-CL-CURRENT: [546/281.7](#); [548/203](#), [548/236](#), [548/311.1](#), [548/465](#), [549/266](#)

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KWIC	Draw. De
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☐ 15. Document ID: US 20030096381 A1

L7: Entry 15 of 40

File: PGPB

May 22, 2003

PGPUB-DOCUMENT-NUMBER: 20030096381

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20030096381 A1

TITLE: Production of polyketides

PUBLICATION-DATE: May 22, 2003

## INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Julien, Bryan	Oakland	CA	US	
Katz, Leonard	Hayward	CA	US	
Khosla, Chaitan	Palo Alto	CA	US	

US-CL-CURRENT: [435/118](#); [435/193](#), [435/252.3](#), [435/320.1](#), [435/69.1](#), [536/23.2](#)

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KWIC	Draw. De
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☐ 16. Document ID: US 20030073205 A1

L7: Entry 16 of 40

File: PGPB

Apr 17, 2003

PGPUB-DOCUMENT-NUMBER: 20030073205

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20030073205 A1

TITLE: Production of polyketides

PUBLICATION-DATE: April 17, 2003

## INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Arslanian, Robert L.	Pacifica	CA	US	
Ashley, Gary	Alameda	CA	US	
Frykman, Scott	Hayward	CA	US	
Julien, Bryan	Oakland	CA	US	
Katz, Leonard	Oakland	CA	US	
Khosla, Chaitan	Palo Alto	CA	US	
Lau, Janice	San Mateo	CA	US	
Licari, Peter J.	Fremont	CA	US	
Regentin, Rika	Hayward	CA	US	
Santi, Daniel	San Francisco	CA	US	
Tang, Li	Foster City	CA	US	

US-CL-CURRENT: 435/117; 435/123, 435/252.3, 544/333, 546/148, 546/167, 546/281.7, 548/204

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KMC	Draw. De
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☐ 17. Document ID: US 20030069277 A1

L7: Entry 17 of 40

File: PGPB

Apr 10, 2003

PGPUB-DOCUMENT-NUMBER: 20030069277

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20030069277 A1

TITLE: Synthesis of epothilones, intermediates thereto and analogues thereof

PUBLICATION-DATE: April 10, 2003

## INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Danishefsky, Samuel J.	Englewood	NJ	US	
Bertinato, Peter	Old Lyme	CT	US	
Su, Dai-Shi	Ambler	PA	US	
Meng, Dongfang	New York	NY	US	
Chou, Ting-Chao	Paramus	NJ	US	
Kamenecka, Ted	New Brunswick	NJ	US	
Sorensen, Erik J.	San Diego	CA	US	
Balog, Aaron	New York	NY	US	
Savin, Kenneth A.	Indianapolis	IN	US	
Kuduk, Scott	Harleysville	PA	US	
Harris, Christina	New York	NY	US	
Zhang, Xiu-Guo	New York	NY	US	
Bertino, Joseph R.	Branford	CT	US	

US-CL-CURRENT: [514/336](#); [514/365](#), [514/397](#), [514/414](#), [514/450](#), [546/281.7](#), [548/204](#),  
[548/236](#), [548/311.1](#), [548/465](#), [549/270](#)

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KWIC	Draw. De
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☐ 18. Document ID: US 20030045711 A1

L7: Entry 18 of 40

File: PGPB

Mar 6, 2003

PGPUB-DOCUMENT-NUMBER: 20030045711  
PGPUB-FILING-TYPE: new  
DOCUMENT-IDENTIFIER: US 20030045711 A1

TITLE: Epothilone derivatives and methods for making and using the same

PUBLICATION-DATE: March 6, 2003

INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Ashley, Gary	Alameda	CA	US	
Arsanian, Robert L.	Pacifica	CA	US	
Carney, John	San Bruno	CA	US	
Metcalfe, Brian	Moraga	CA	US	
Tang, Li	Foster City	CA	US	

US-CL-CURRENT: [540/462](#); [549/269](#)

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KWIC	Draw. De
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☐ 19. Document ID: US 20030023082 A1

L7: Entry 19 of 40

File: PGPB

Jan 30, 2003

PGPUB-DOCUMENT-NUMBER: 20030023082  
PGPUB-FILING-TYPE: new  
DOCUMENT-IDENTIFIER: US 20030023082 A1

TITLE: Epothilone derivatives and methods for making and using the same

PUBLICATION-DATE: January 30, 2003

INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Ashley, Gary	Alameda	CA	US	
Metcalfe, Brian	Moraga	CA	US	

US-CL-CURRENT: [540/461](#); [544/278](#), [546/115](#), [548/207](#), [548/242](#), [548/360.5](#)

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KWIC	Draw. De
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☐ 20. Document ID: US 20020193423 A1

L7: Entry 20 of 40

File: PGPB

Dec 19, 2002

PGPUB-DOCUMENT-NUMBER: 20020193423

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20020193423 A1

TITLE: Bioactive compound

PUBLICATION-DATE: December 19, 2002

## INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Northcote, Peter T.	Wellington		NZ	
Miller, John H.	Wellington		NZ	
Hood, Kylie A.	Wellington		NZ	
West, Lyndon M.	Wellington		NZ	

US-CL-CURRENT: 514/450; 549/267

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KWIC	Draw. De
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**Search Results - Record(s) 21 through 40 of 40 returned.**☐ 21. Document ID: US 20020193361 A1**Using default format because multiple data bases are involved.**

L7: Entry 21 of 40

File: PGPB

Dec 19, 2002

PGPUB-DOCUMENT-NUMBER: 20020193361

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20020193361 A1

TITLE: Epothilone derivatives and methods for making and using the same

PUBLICATION-DATE: December 19, 2002

## INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Ashley, Gary	Alameda	CA	US	
Metcalf, Brian	Moraga	CA	US	

US-CL-CURRENT: 514/183; 514/450, 540/461, 549/269

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KWIC	Draw. Data
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☐ 22. Document ID: US 20020156110 A1

L7: Entry 22 of 40

File: PGPB

Oct 24, 2002

PGPUB-DOCUMENT-NUMBER: 20020156110

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20020156110 A1

TITLE: Epothilone compounds and methods for making and using the same

PUBLICATION-DATE: October 24, 2002

## INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Arslanian, Robert L.	Pacifica	CA	US	
Carney, John R.	San Bruno	CA	US	
Metcalf, Brian	Moraga	CA	US	

US-CL-CURRENT: 514/365; 548/204

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KWIC	Draw. De
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☐ 23. Document ID: US 20020137152 A1

L7: Entry 23 of 40

File: PGPB

Sep 26, 2002

PGPUB-DOCUMENT-NUMBER: 20020137152  
PGPUB-FILING-TYPE: new  
DOCUMENT-IDENTIFIER: US 20020137152 A1

TITLE: Fermentation process for epothilones

PUBLICATION-DATE: September 26, 2002

## INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Santi, Daniel	San Francisco	CA	US	
Metcalfe, Brian	Moraga	CA	US	
Ashley, Gary	Alameda	CA	US	

US-CL-CURRENT: 435/118; 435/184, 435/252.3

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KWIC	Draw. De
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☐ 24. Document ID: US 20020062030 A1

L7: Entry 24 of 40

File: PGPB

May 23, 2002

PGPUB-DOCUMENT-NUMBER: 20020062030  
PGPUB-FILING-TYPE: new  
DOCUMENT-IDENTIFIER: US 20020062030 A1

TITLE: Method for synthesizing epothilones and epothilone analogs

PUBLICATION-DATE: May 23, 2002

## INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
White, James David	Philomath	OR	US	
Carter, Rich Garrett	Oxford	MS	US	
Sundermann, Kurt Frederick	Corvallis	OR	US	

US-CL-CURRENT: 548/204

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KWIC	Draw. De
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☐ 25. Document ID: US 20020058817 A1



L7: Entry 25 of 40

File: PGPB

May 16, 2002

PGPUB-DOCUMENT-NUMBER: 20020058817  
PGPUB-FILING-TYPE: new  
DOCUMENT-IDENTIFIER: US 20020058817 A1

TITLE: Synthesis of epothilones, intermediates thereto and analogues thereof

PUBLICATION-DATE: May 16, 2002

## INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Danishefsky, Samuel J.	Englewood	NJ	US	
Stachel, Shawn J.	Perkasie	PA	US	
Lee, Chul Bom	Princeton	NJ	US	
Chappell, Mark D.	Noblesville	IN	US	
Wu, Zhicai	New York	NY	US	

US-CL-CURRENT: 546/281.7; 540/451, 548/194, 548/236, 548/311.1, 548/466, 549/266

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KWIC	Draw D
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☐ 26. Document ID: US 20020058286 A1

L7: Entry 26 of 40

File: PGPB

May 16, 2002

PGPUB-DOCUMENT-NUMBER: 20020058286  
PGPUB-FILING-TYPE: new  
DOCUMENT-IDENTIFIER: US 20020058286 A1

TITLE: Synthesis of epothilones, intermediates thereto and analogues thereof

PUBLICATION-DATE: May 16, 2002

## INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Danishefsky, Samuel J.	Englewood	NJ	US	
Stachel, Shawn J.	Perkasi	PA	US	
Lee, Chul Bom	Princeton	NJ	US	
Chappell, Mark D.	Noblesville	IN	US	
Chou, Ting-Chao	Paramus	NJ	US	
Wu, Zhicai	New York	NY	US	

US-CL-CURRENT: 435/7.1; 534/10, 536/17.4, 540/451, 549/266

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KWIC	Draw D
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☐ 27. Document ID: US 20020045609 A1

L7: Entry 27 of 40

File: PGPB

Apr 18, 2002

PGPUB-DOCUMENT-NUMBER: 20020045609  
PGPUB-FILING-TYPE: new  
DOCUMENT-IDENTIFIER: US 20020045609 A1

TITLE: Epothilone derivatives and methods for making and using the same

PUBLICATION-DATE: April 18, 2002

## INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Ashley, Gary	Alameda	CA	US	
Fardis, Maria	San Carlos	CA	US	
Santi, Daniel	San Francisco	CA	US	

US-CL-CURRENT: 514/183; 514/450, 540/451, 549/270

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KWIC	Draw. De
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☐ 28. Document ID: US 6683100 B2

L7: Entry 28 of 40

File: USPT

Jan 27, 2004

US-PAT-NO: 6683100  
DOCUMENT-IDENTIFIER: US 6683100 B2

TITLE: Organic compounds

DATE-ISSUED: January 27, 2004

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
van Hoogevest; Peter	Bubendorf			CH

US-CL-CURRENT: 514/365; 514/456

## ABSTRACT:

Pharmaceutical formulations comprising an epothilone in the form of an infusion concentrate or a lyophilised composition, and methods of administration of an epothilone in suitable form for parenteral administration.

20 Claims, 0 Drawing figures  
Exemplary Claim Number: 1

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KWIC	Draw. De
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☐ 29. Document ID: US 6660758 B1

L7: Entry 29 of 40

File: USPT

Dec 9, 2003

US-PAT-NO: 6660758

DOCUMENT-IDENTIFIER: US 6660758 B1

TITLE: Epothilone analogs

DATE-ISSUED: December 9, 2003

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Nicolaou; Kyriacos C.	La Jolla	CA		
He; Yun	San Diego	CA		
Ninkovic; Sacha	San Diego	CA		
Pastor; Joaquin	Madrid			ES
Roschangar; Frank	Durham	NC		
Sarabia; Francisco	Torre De Benagalbon			ES
Vallberg; Hans	Huddinge			SE
Vourloumis; Dionisios	Apex	NC		
Winssinger; Nicolas	La Jolla	CA		
Yang; Zhen	Brookline	MA		
King; Nigel Paul	Camborne			GB
Finlay; M. Ray	Killinchy			GB

US-CL-CURRENT: 514/374; 548/235

## ABSTRACT:

Epothilone A, epothilone B, analogs of epothilone and libraries of epothilone analogs are synthesized. Epothilone A and B are known anticancer agents that derive their anticancer activity by the prevention of mitosis through the induction and stabilization of microtubule assembly. The analogs of epothilone are novel. Several of the analogs are demonstrated to have a superior cytotoxic activity as compared to epothilone A or epothilone B as demonstrated by their enhanced ability to induce the polymerization and stabilization of microtubules.

13 Claims, 101 Drawing figures

Exemplary Claim Number: 1

Number of Drawing Sheets: 101

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWIC	Draw. De
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☐ 30. Document ID: US 6603023 B2

L7: Entry 30 of 40

File: USPT

Aug 5, 2003

US-PAT-NO: 6603023

DOCUMENT-IDENTIFIER: US 6603023 B2

TITLE: Synthesis of epothilones, intermediates thereto and analogues thereof

DATE-ISSUED: August 5, 2003

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Danishefsky; Samuel J.	Englewood	NJ		
Bertinato; Peter	Old Lyme	CT		
Su; Dai-Shi	Ambler	PA		
Meng; DongFang	New York	NY		
Chou; Ting-Chao	Paramus	NJ		
Kamenecka; Ted	New Brunswick	NJ		
Sorensen; Erik J.	San Diego	CA		
Balog; Aaron	New York	NY		
Savin; Kenneth A.	Indianapolis	IN		
Kuduk; Scott	Harleysville	PA		
Harris; Christina	New York	NY		
Zhang; Xiu-Guo	New York	NY		
Bertino; Joseph R.	Branford	CT		

US-CL-CURRENT: 549/346

## ABSTRACT:

The present invention provides convergent processes for preparing epothilone A and B, desoxyepothilones A and B, and analogues thereof, useful in the treatment of cancer and cancer which has developed a multidrug-resistant phenotype. Also provided are intermediates useful for preparing said epothilones.

23 Claims, 117 Drawing figures

Exemplary Claim Number: 1

Number of Drawing Sheets: 102

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWC	Draw. De
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☐ 31. Document ID: US 6596875 B2

L7: Entry 31 of 40

File: USPT

Jul 22, 2003

US-PAT-NO: 6596875

DOCUMENT-IDENTIFIER: US 6596875 B2

TITLE: Method for synthesizing epothilones and epothilone analogs

DATE-ISSUED: July 22, 2003

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
White; James David	Philomath	OR	97370	
Carter; Rich Garrett	Oxford	MS	38655	
Sundermann; Kurt Frederick	Corvallis	OR	97339	

US-CL-CURRENT: 548/204

## ABSTRACT:

A method for making epothilones and epothilone analogs is described, as are novel compounds made by the method. One embodiment of the method was used to synthesize epothilone B by a convergent approach that entailed Wittig coupling of an ylide derived from phosphonium bromide with an aldehyde. The former was prepared from propargyl alcohol by a nine-step pathway which installed both trisubstituted double bonds with clean Z configuration. Macrolactonization of a resulting seco acid provided the following intermediate diene epothilone analog. Selective saturation of the 9,10-olefin and subsequent epoxidation provided epothilone B. ##STR1##

30 Claims, 0 Drawing figures

Exemplary Claim Number: 1

Full	Title	Citation	Front	Review	Classification	Date	Reference			Claims	KnowlC	Draw De
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☐ 32. Document ID: US 6589968 B2

L7: Entry 32 of 40

File: USPT

Jul 8, 2003

US-PAT-NO: 6589968

DOCUMENT-IDENTIFIER: US 6589968 B2

TITLE: Epothilone compounds and methods for making and using the same

DATE-ISSUED: July 8, 2003

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Arsanian; Robert L.	Pacifica	CA		
Carney; John R.	San Bruno	CA		
Metcalf; Brian	Moraga	CA		

US-CL-CURRENT: 514/365; 548/204

## ABSTRACT:

This present invention relates to compounds of formula (I) ##STR1##

and to pharmaceutically acceptable salts and solvates thereof, wherein R.sup.1, R.sup.2, R.sup.3, R.sup.4, R.sup.5, W, X, Y, and Ar are as defined herein. Compounds of formula (I) are useful in the treatment of diseases or conditions characterized by cellular hyperproliferation. This invention also relates to means for the preparation of compounds of formula (I); formulations containing compounds of formula (I); and methods for the use of said compounds and formulations in the treatment of a disease or condition characterized by cellular hyperproliferation, including cancer.

8 Claims, 0 Drawing figures

Exemplary Claim Number: 1

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWC	Draw. De
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☐ 33. Document ID: US 6583290 B1

L7: Entry 33 of 40

File: USPT

Jun 24, 2003

US-PAT-NO: 6583290

DOCUMENT-IDENTIFIER: US 6583290 B1

TITLE: 14-methyl epothilone derivatives

DATE-ISSUED: June 24, 2003

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Julien; Bryan	Oakland	CA		
Katz; Leonard	Hayward	CA		
Khosla; Chaitan	Palo Alto	CA		
Tang; Li	Foster City	CA		
Ziermann; Rainer	San Mateo	CA		

US-CL-CURRENT: 548/203; 181/205, 546/268.1

## ABSTRACT:

Compounds of the invention include 14-mehtyl epothilone derivatives. More generally, preferred compounds of the invention are those that can be produced by altering the epothilone PKS genes as described herein and optionally by action of epothilone modification enzymes and/or by chemically modifying the resulting epothilones produces when those genes are expressed.

2 Claims, 9 Drawing figures

Exemplary Claim Number: 1

Number of Drawing Sheets: 8

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWC	Draw. De
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☐ 34. Document ID: US 6489314 B1

L7: Entry 34 of 40

File: USPT

Dec 3, 2002

US-PAT-NO: 6489314

DOCUMENT-IDENTIFIER: US 6489314 B1

TITLE: Epothilone derivatives and methods for making and using the same

DATE-ISSUED: December 3, 2002

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Ashley; Gary	Alameda	CA		
Metcalf; Brian	Moraga	CA		

US-CL-CURRENT: 514/183; 540/451, 540/455, 540/461, 540/462, 540/463

## ABSTRACT:

The present invention relates to 16-membered macrocyclic compounds. In one aspect of the present invention, compounds of the formula ##STR1##

are provided wherein: R.sup.1, R.sup.2, R.sup.3, and R.sup.5 are each independently hydrogen, C.sub.1 -C.sub.10 alkyl, C.sub.2 -C.sub.10 alkenyl, C.sub.2 -C.sub.10 alkynyl, aryl or alkylaryl; R.sup.4 is hydrogen, halogen, C.sub.1 -C.sub.10 alkyl, C.sub.1 -C.sub.10 hydroxyalkyl, C.sub.1 -C.sub.10 haloalkyl, aryl, --C(.dbd.O)R.sup.6, --C(.dbd.O)OR.sup.6, --NR.sup.6 R.sup.7 where R.sup.6 and R.sup.7 are each independently hydrogen, C.sub.1 -C.sub.10 aliphatic, aryl or alkylaryl; W is O, NR.sup.8 where R.sup.8 is hydrogen, C.sub.1 -C.sub.10 alkyl, C.sub.2 -C.sub.10 alkenyl, C.sub.2 -C.sub.10 alkynyl, aryl or alkylaryl; X is O, CH.sub.2 or a carbon-carbon double bond; Y is absent or a C.sub.1 -C.sub.10 alkyl, C.sub.2 -C.sub.10 alkenyl, or C.sub.2 -C.sub.10 alkynyl; and Ar is aryl; provided that 10,11-dehydroepothilone C is excluded.

27 Claims, 0 Drawing figures

Exemplary Claim Number: 1

Full	Title	Citation	Front	Review	Classification	Date	Reference	Abstract	Claims	KWIC	Draw. De
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☐ 35. Document ID: US 6441186 B1

L7: Entry 35 of 40

File: USPT

Aug 27, 2002

US-PAT-NO: 6441186

DOCUMENT-IDENTIFIER: US 6441186 B1

TITLE: Epothilone analogs

DATE-ISSUED: August 27, 2002

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Nicolaou; Kyriacos C.	La Jolla	CA		
He; Yun	San Diego	CA		
Ninkovic; Sacha	San Diego	CA		
Pastor; Joaquin	San Diego	CA		
Roschangar; Frank	San Diego	CA		
Sarabia; Francisco	Torre de Benagalbon			ES
Vallberg; Hans	Huddinge			SE
Vourloumis; Dionisios	San Diego	CA		
Winssinger; Nicolas	La Jolla	CA		
Yang; Zhen	San Diego	CA		

King; N. Paul                      San Diego                      CA  
Finlay; M. Ray                      San Diego                      CA

US-CL-CURRENT: 548/204

ABSTRACT:

Epothilone A, epothilone B, analogs of epothilone and libraries of epothilone analogs are synthesized. Epothilone A and B are known anticancer agents that derive their anticancer activity by the prevention of mitosis through the induction and stabilization of microtubulin assembly. The analogs of epothilone are novel. Several of the anlogs are demonstrated to have a superior cytotoxic activities as compared to epothilone A or epothilone B as demonstrated by their enhanced ability to induce the polymerization and stabilization of microtubules.

1 Claims, 74 Drawing figures  
Exemplary Claim Number: 1  
Number of Drawing Sheets: 74

Full	Title	Citation	Front	Review	Classification	Date	Reference	Abstract	Claims	KWIC	Draw De
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☐ 36. Document ID: US 6410301 B1

L7: Entry 36 of 40

File: USPT

Jun 25, 2002

US-PAT-NO: 6410301

DOCUMENT-IDENTIFIER: US 6410301 B1

TITLE: Myxococcus host cells for the production of epothilones

DATE-ISSUED: June 25, 2002

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Julien; Bryan	Oakland	CA		
Katz; Leonard	Hayward	CA		
Khosla; Chaitan	Palo Alto	CA		

US-CL-CURRENT: 435/252.3

ABSTRACT:

Recombinant Myxococcus host cell containing recombinant expression vectors containing epothilone polyketide synthase genes can produce epothilones C and D.

6 Claims, 4 Drawing figures  
Exemplary Claim Number: 1  
Number of Drawing Sheets: 4

Full	Title	Citation	Front	Review	Classification	Date	Reference	Abstract	Claims	KWIC	Draw De
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☐ 37. Document ID: US 6316630 B1

L7: Entry 37 of 40

File: USPT

Nov 13, 2001

US-PAT-NO: 6316630

DOCUMENT-IDENTIFIER: US 6316630 B1

TITLE: Synthesis of epothilones, intermediates thereto and analogues thereof

DATE-ISSUED: November 13, 2001

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Danishefsky; Samuel J.	Englewood	NJ		
Bertino; Peter	Old Lyme	CT		
Su; Dai-Shi	New York	NY		
Meng; DongFang	New York	NY		
Chou; Ting-Chao	Paramus	NJ		
Kamenecka; Ted	New York	NY		
Sorensen; Erik J	San Diego	CA		
Balog; Aaron	New York	NY		
Savin; Kenneth A.	New York	NY		

US-CL-CURRENT: 546/281.7; 546/340, 548/204, 548/510, 549/494, 549/498, 560/174

## ABSTRACT:

The present invention provides convergent processes for preparing epothilone A and B, desoxyepothilones A and B, and analogues thereof, useful in the treatment of cancer and cancer which has developed a multidrug-resistant phenotype. Also provided are intermediates useful for preparing said epothilones.

8 Claims, 75 Drawing figures

Exemplary Claim Number: 1

Number of Drawing Sheets: 102

Full	Title	Citation	Front	Review	Classification	Date	Reference			Claims	FIGS	Draw. Ds
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☐ 38. Document ID: US 6303342 B1

L7: Entry 38 of 40

File: USPT

Oct 16, 2001

US-PAT-NO: 6303342

DOCUMENT-IDENTIFIER: US 6303342 B1

TITLE: Recombinant methods and materials for producing epothilones C and D

DATE-ISSUED: October 16, 2001

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Julien; Bryan	Oakland	CA		
Katz; Leonard	Hayward	CA		
Khosla; Chaitan	Palo Alto	CA		
Tang; Li	Foster City	CA		

US-CL-CURRENT: 435/76

## ABSTRACT:

Recombinant nucleic acids that encode all or a portion of the epothilone polyketide synthase (PKS) are used to express recombinant PKS genes in host cells for the production of epothilones, epothilone derivatives, and polyketides that are useful as cancer chemotherapeutics, fungicides, and immunosuppressants.

29 Claims, 9 Drawing figures  
Exemplary Claim Number: 1  
Number of Drawing Sheets: 8

Full	Title	Citation	Front	Review	Classification	Date	Reference	Abstract	Claims	KOMC	Draw De
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☐ 39. Document ID: US 6302838 B1

L7: Entry 39 of 40

File: USPT

Oct 16, 2001

US-PAT-NO: 6302838

DOCUMENT-IDENTIFIER: US 6302838 B1

**\*\* See image for Certificate of Correction \*\***

TITLE: Cancer treatment with epothilones

DATE-ISSUED: October 16, 2001

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
O'Reilly; Terence	Basel			CH
Wartmann; Markus	Riehen			CH
Litchman; Manuel	Teaneck	NJ		
Cohen; Pamela	Tenafly	NJ		

US-CL-CURRENT: 514/365

## ABSTRACT:

The invention relates to the treatment of a proliferative disease, especially according to certain treatment regimens, with an epothilone, especially with epothilone A and more preferably epothilone B; as well as to the treatment of certain cancers with such an epothilone.

24 Claims, 0 Drawing figures  
Exemplary Claim Number: 1

Full	Title	Citation	Front	Review	Classification	Date	Reference	Abstract	Claims	Drawings	Draw. De
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☐ 40. Document ID: US 6204388 B1

L7: Entry 40 of 40

File: USPT

Mar 20, 2001

US-PAT-NO: 6204388

DOCUMENT-IDENTIFIER: US 6204388 B1

TITLE: Synthesis of epothilones, intermediates thereto and analogues thereof

DATE-ISSUED: March 20, 2001

## INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Danishefsky; Samuel J.	Englewood	NJ		
Bertinato; Peter	Old Lyme	CT		
Su; Dai-Shi	New York	NY		
Meng; DongFang	New York	NY		
Chou; Ting-Chao	Paramus	NJ		
Kamenecka; Ted	New York	NY		
Sorensen; Erik J	San Diego	CA		
Balog; Aaron	New York	NY		
Savin; Kenneth A.	New York	NY		
Kuduk; Scott	Harleysville	PA		
Harris; Christina	New York	NY		
Zhang; Xiu-Guo	New York	NY		
Bertino; Joseph R.	Branford	CT		

US-CL-CURRENT: 546/340; 548/204, 548/510, 549/494, 549/498, 560/174

## ABSTRACT:

The present invention provides convergent processes for preparing epothilone A and B, desoxyepothilones A and B, and analogues thereof, useful in the treatment of cancer and cancer which has developed a multidrug-resistant phenotype. Also provided are intermediates useful for preparing said epothilones.

9 Claims, 117 Drawing figures

Exemplary Claim Number: 1

Number of Drawing Sheets: 102

Full	Title	Citation	Front	Review	Classification	Date	Reference	Abstract	Claims	Drawings	Draw. De
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Clear	Generate Collection	Print	Fwd Refs	Bkwd Refs	Generate OACS
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Terms	Documents
L6 and cellulosum	40

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## STN SEARCH

09/724,876

5/11/04

=&gt; file .nash

=&gt; s desoxyepothilone

L1 18 FILE MEDLINE  
L2 48 FILE CAPLUS  
L3 52 FILE SCISEARCH  
L4 4 FILE LIFESCI  
L5 24 FILE BIOSIS  
L6 25 FILE EMBASE

TOTAL FOR ALL FILES

L7 171 DESOXYEPOTHILONE

=&gt; s epothilone d

L8 20 FILE MEDLINE  
L9 99 FILE CAPLUS  
L10 21 FILE SCISEARCH  
L11 4 FILE LIFESCI  
L12 26 FILE BIOSIS  
L13 37 FILE EMBASE

TOTAL FOR ALL FILES

L14 207 EPOTHILONE D

=&gt; d

L14 ANSWER 1 OF 207 MEDLINE on STN  
AN 2003604225 MEDLINE  
DN PubMed ID: 14686728  
TI Nanomolar concentrations of **epothilone D** inhibit the  
proliferation of glioma cells and severely affect their tubulin  
cytoskeleton.  
AU Dietzmann A; Kanakis D; Kirches E; Kropf S; Mawrin C; Dietzmann K  
CS Department of Neuropathology, University of Magdeburg, Magdeburg, Germany.  
SO Journal of neuro-oncology, (2003 Nov) 65 (2) 99-106.  
Journal code: 8309335. ISSN: 0167-594X.  
CY Netherlands  
DT Journal; Article; (JOURNAL ARTICLE)  
LA English  
FS Priority Journals  
EM 200402  
ED Entered STN: 20031223  
Last Updated on STN: 20040203  
Entered Medline: 20040202

=&gt; s l7 or l14

L15 37 FILE MEDLINE  
L16 142 FILE CAPLUS  
L17 69 FILE SCISEARCH  
L18 8 FILE LIFESCI  
L19 50 FILE BIOSIS  
L20 59 FILE EMBASE

TOTAL FOR ALL FILES

L21 365 L7 OR L14

=&gt; s l21 not 2000-2004/py

L22 3 FILE MEDLINE  
L23 26 FILE CAPLUS  
L24 5 FILE SCISEARCH  
L25 2 FILE LIFESCI  
L26 4 FILE BIOSIS  
L27 6 FILE EMBASE

TOTAL FOR ALL FILES

L28 46 L21 NOT 2000-2004/PY

=&gt; s l28 and biosynthesis

L29 0 FILE MEDLINE  
L30 0 FILE CAPLUS

L31 0 FILE SCISEARCH  
L32 0 FILE LIFESCI  
L33 0 FILE BIOSIS  
L34 0 FILE EMBASE

TOTAL FOR ALL FILES

L35 0 L28 AND BIOSYNTHESIS

=> dup rem 128

PROCESSING COMPLETED FOR L28

L36 28 DUP REM L28 (18 DUPLICATES REMOVED)

=> d ibib ab 1-28

L36 ANSWER 1 OF 28 CAPLUS COPYRIGHT 2004 ACS on STN  
ACCESSION NUMBER: 2000:514331 CAPLUS  
DOCUMENT NUMBER: 134:100671  
TITLE: Chapter I: The first total syntheses of epothilones A,  
B, C and D. Chapter II: The first total syntheses of  
12-epi-CP-263,114, and 12-epi-CP-225,917  
AUTHOR(S): Meng, Dongfang  
CORPORATE SOURCE: Columbia University, USA  
SOURCE: (1999) 326 pp. Avail.: University Microfilms  
International, Order No. DA9949022  
From: Diss. Abstr. Int., B 2000, 60(10), 5096  
DOCUMENT TYPE: Dissertation  
LANGUAGE: English  
AB Unavailable

L36 ANSWER 2 OF 28 CAPLUS COPYRIGHT 2004 ACS on STN DUPLICATE 1  
ACCESSION NUMBER: 1999:680932 CAPLUS  
DOCUMENT NUMBER: 132:22767  
TITLE: Complex Target-Oriented Synthesis in the Drug  
Discovery Process: A Case History in the dEpoB Series  
AUTHOR(S): Harris, Christina R.; Danishefsky, Samuel J.  
CORPORATE SOURCE: Laboratory for Bioorganic Chemistry, Sloan-Kettering  
Institute for Cancer Research, New York, NY, 10021,  
USA  
SOURCE: Journal of Organic Chemistry (1999), 64(23), 8434-8456  
CODEN: JOCEAH; ISSN: 0022-3263  
PUBLISHER: American Chemical Society  
DOCUMENT TYPE: Journal; General Review  
LANGUAGE: English  
AB A review with 103 refs. on complex target-oriented synthesis of naturally  
occurring cytotoxic agents of potential clin. value in the chemotherapy of  
cancer. In particular, the crit. role of complex target-oriented  
synthesis in the discovery process pertinent to 12,13-  
**desoxyepothilone** B is described.  
REFERENCE COUNT: 154 THERE ARE 154 CITED REFERENCES AVAILABLE FOR  
THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE  
FORMAT

L36 ANSWER 3 OF 28 CAPLUS COPYRIGHT 2004 ACS on STN DUPLICATE 2  
ACCESSION NUMBER: 1999:444724 CAPLUS  
DOCUMENT NUMBER: 131:286299  
TITLE: New Chemical Synthesis of the Promising Cancer  
Chemotherapeutic Agent 12,13-**Desoxyepothilone**  
B: Discovery of a Surprising Long-Range Effect on the  
Diastereoselectivity of an Aldol Condensation  
AUTHOR(S): Harris, Christina R.; Kuduk, Scott D.; Balog, Aaron;  
Savin, Ken; Glunz, Peter W.; Danishefsky, Samuel J.  
CORPORATE SOURCE: Laboratory for Bioorganic Chemistry, The  
Sloan-Kettering Institute for Cancer Research, New  
York, NY, 10021, USA  
SOURCE: Journal of the American Chemical Society (1999),  
121(30), 7050-7062  
CODEN: JACSAT; ISSN: 0002-7863  
PUBLISHER: American Chemical Society  
DOCUMENT TYPE: Journal  
LANGUAGE: English  
OTHER SOURCE(S): CASREACT 131:286299

AB The epothilones are naturally occurring cytotoxic mols. that possess the remarkable ability to arrest cell division through the stabilization of microtubule assemblies. In vivo studies with 12,13-**desoxyepothilone B** (dEpoB) (I), have established that the desoxy compd. is well tolerated and virtually curative against a variety of sensitive and resistant xenograft tumors in animal models. In light of these discoveries, a chem. synthesis of dEpoB would be able to support a serious and substantial discovery research program directed toward the clin. development of this mol. The overall strategy for this endeavor assumed the ability to synthesize dEpoB from three constructs which include an achiral .beta.,.delta.-diketo ester construct A (II), an (S)-2-methylpentenal moiety B (III), and the thiazoyl-contg. vinyl iodide moiety C (IV). It was envisioned that a diastereoselective aldol condensation between an achiral C5-C6 (Z)-metalloenolate derived from construct A and an (S)-2-methylalkanal fragment, B, would generate the desired C6-C7 bond. Second, a B-alkyl Suzuki coupling between the vinyl iodide construct C and an alkyl borane would form the C11-C12 bond. Finally, a late-stage redn. of the C3 ketone to the requisite C3 alc. with high asym. induction would permit introduction of the .beta.,.delta.-diketo ester fragment A, into the synthesis as a readily accessible achiral building block. The governing concepts the new synthesis are described.

REFERENCE COUNT: 12 THERE ARE 12 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L36 ANSWER 4 OF 28 CAPLUS COPYRIGHT 2004 ACS on STN DUPLICATE 3

ACCESSION NUMBER: 1999:589150 CAPLUS

DOCUMENT NUMBER: 131:336853

TITLE: The synthesis and evaluation of 12,13-

AUTHOR(S): benzodesoxyepothilone B: a highly convergent route  
Glunz, Peter W.; He, Lifeng; Horwitz, Susan B.;  
Chakravarty, Subrata; Ojima, Iwao; Chou, Ting-Chao;  
Danishefsky, Samuel J.

CORPORATE SOURCE: Laboratory for Bioorganic Chemistry, Sloan-Kettering  
Institute for Cancer Research, New York, NY, 10021,  
USA

SOURCE: Tetrahedron Letters (1999), 40(38), 6895-6898

CODEN: TELEAY; ISSN: 0040-4039

PUBLISHER: Elsevier Science Ltd.

DOCUMENT TYPE: Journal

LANGUAGE: English

OTHER SOURCE(S): CASREACT 131:336853

AB The title compd. I retains some of the affinity for microtubule assemblies as does 12,13-**desoxyepothilone B** (II).

REFERENCE COUNT: 19 THERE ARE 19 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L36 ANSWER 5 OF 28 CAPLUS COPYRIGHT 2004 ACS on STN DUPLICATE 4

ACCESSION NUMBER: 1999:183184 CAPLUS

DOCUMENT NUMBER: 130:352108

TITLE: Remarkable long range effects on the diastereoface  
selectivity in an aldol condensation

AUTHOR(S): Harris, Christina R.; Kuduk, Scott D.; Balog, Aaron;  
Savin, Ken A.; Danishefsky, Samuel J.

CORPORATE SOURCE: Laboratory Bioorganic Chemistry, Sloan-Kettering  
Institute Cancer Research, New York, NY, 10021, USA

SOURCE: Tetrahedron Letters (1999), 40(12), 2267-2270

CODEN: TELEAY; ISSN: 0040-4039

PUBLISHER: Elsevier Science Ltd.

DOCUMENT TYPE: Journal

LANGUAGE: English

AB The stereochem. results in an aldol reaction of 12,13-**desoxyepothilone B** between the Li enolates of I (R = Me, Et3Si) and various .alpha.-Me aldehydes indicates a stabilizing through space interaction between C4-C5 unsatn. and the formyl group. This interaction leads to a reaction conformation which favors a C7-C8 (epothilone numbering) anti-relationship in the aldol products. Included is an extensive study that identifies steric and electronic effects of various .alpha.-Me aldehydes in the aldol diastereoselection.

REFERENCE COUNT: 7 THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L36 ANSWER 6 OF 28 CAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 1999:183183 CAPLUS  
DOCUMENT NUMBER: 130:337931  
TITLE: Dianion equivalents corresponding to the  
polypropionate domain of epothilone B  
AUTHOR(S): Harris, Christina R.; Kuduk, Scott D.; Savin, Ken;  
Balog, Aaron; Danishefsky, Samuel J.  
CORPORATE SOURCE: Laboratory Bioorganic Chemistry, Sloan-Kettering  
Institute Cancer Research, New York, NY, 10021, USA  
SOURCE: Tetrahedron Letters (1999), 40(12), 2263-2266  
CODEN: TELEAY; ISSN: 0040-4039  
PUBLISHER: Elsevier Science Ltd.  
DOCUMENT TYPE: Journal  
LANGUAGE: English  
OTHER SOURCE(S): CASREACT 130:337931  
AB A modified synthesis of the polypropionate portion of epothilone, which  
utilizes a novel, diastereoselective aldol reaction of  
(S)-2-methyl-4-pentenol and the Z-enolate of the tricarboxyl species  
EtCOCMe2COCH2CO2CMe3 is reported.  
REFERENCE COUNT: 10 THERE ARE 10 CITED REFERENCES AVAILABLE FOR THIS  
RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L36 ANSWER 7 OF 28 CAPLUS COPYRIGHT 2004 ACS on STN DUPLICATE 5

ACCESSION NUMBER: 2000:555968 CAPLUS  
DOCUMENT NUMBER: 133:275843  
TITLE: Epothilone A-D and their thiazole-modified analogs as  
novel anticancer agents  
AUTHOR(S): Hofle, G.; Glaser, N.; Leibold, T.; Sefkow, M.  
CORPORATE SOURCE: Dep. Nat. Product Chem., GBF, Gesellschaft Biotechnol.  
Forschung mbH, Braunschweig, D-38124, Germany  
SOURCE: Pure and Applied Chemistry (1999), 71(11), 2019-2024  
CODEN: PACHAS; ISSN: 0033-4545  
PUBLISHER: Blackwell Science Ltd.  
DOCUMENT TYPE: Journal  
LANGUAGE: English  
AB Starting from epothilone A-D obtained by large scale fermn. of the  
myxobacterium Sorangium cellulosum, the thiazole side-chain was  
extensively modified by substitution, oxidn. and replacement. Metalation  
afforded the C-19 carbanion which was quenched by various carbon and  
heteroatom electrophiles to give C-19 substituted epothilones. Thiazole  
N-oxides were obtained by treatment of epothilone A and B with  
m-chloroperbenzoic acid and rearranged by acetic anhydride to 21-acetoxy  
epothilones. Cleavage of epothilones A and B with ozone gave Me ketones  
from which carbonyl derivs. and aldol condensation products were prepd.  
Similarly vinyl boronic acid was obtained and transformed by Suzuki  
coupling or iodination/Stille coupling to aryl and heteroaryl analogs.  
The structure-activity relationships for thiazolyl side chain of  
epothilones were in line with published data obtained from analogs prepd.  
by total synthesis. Only few modifications were tolerated without  
significant loss of activity, i.e. replacement of the thiazole by an  
oxazole ring or introduction of small substituents at C-21.  
REFERENCE COUNT: 49 THERE ARE 49 CITED REFERENCES AVAILABLE FOR THIS  
RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L36 ANSWER 8 OF 28 CAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 1999:470234 CAPLUS  
DOCUMENT NUMBER: 131:286303  
TITLE: N-oxidation of epothilone A-C and O-acyl rearrangement  
to C-19- and C-21-substituted epothilones  
AUTHOR(S): Hofle, Gerhard; Glaser, Nicole; Kiffe, Michael; Hecht,  
Hans-Jurgen; Sasse, Florenz; Reichenbach, Hans  
CORPORATE SOURCE: Abteilung Naturstoffchemie Gesellschaft fur  
Biotechnologische Forschung, Braunschweig, D-38124,  
Germany  
SOURCE: Angewandte Chemie, International Edition (1999),  
38(13/14), 1971-1974  
CODEN: ACIEF5; ISSN: 1433-7851  
PUBLISHER: Wiley-VCH Verlag GmbH  
DOCUMENT TYPE: Journal  
LANGUAGE: English



OTHER SOURCE(S): CASREACT 131:286303

AB Epothilones A-C underwent N-oxidn. on treatment with MCPBA in CH<sub>2</sub>Cl<sub>2</sub>. The N-oxide of epothilones A and B were converted to the 2-acetoxymethylthiazole derivs. with Ac<sub>2</sub>O and these were hydrolyzed to epothilones E and F. Some chloro and tosyloxy derivs. were also prepd. In vitro antitumor activities are reported.

REFERENCE COUNT: 35 THERE ARE 35 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L36 ANSWER 9 OF 28 CAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 1999:663080 CAPLUS

DOCUMENT NUMBER: 132:22785

TITLE: Sets of Aldolase Antibodies with Antipodal Reactivities. Formal Synthesis of Epothilone E by Large-Scale Antibody-Catalyzed Resolution of Thiazole Aldol

AUTHOR(S): Sinha, Subhash C.; Sun, Jian; Miller, Gregory; Barbas, Carlos F. III; Lerner, Richard A.

CORPORATE SOURCE: Department of Molecular Biology and the Skaggs Institute for Chemical Biology, The Scripps Research Institute, La Jolla, CA, 92037, USA

SOURCE: Organic Letters (1999), 1(10), 1623-1626

CODEN: ORLEF7; ISSN: 1523-7060

PUBLISHER: American Chemical Society

DOCUMENT TYPE: Journal

LANGUAGE: English

OTHER SOURCE(S): CASREACT 132:22785

AB Three monoclonal aldolase antibodies, generated against a .beta.-diketone hapten by reactive immunization, catalyzed rapid and highly enantioselective retro-aldol reactions of ent-I (R = R<sub>1</sub> = Me; R = Et, Pr, Bu, 1-pentyl, 1-butenyl, CH<sub>2</sub>F, R<sub>1</sub> = Me; R = Me, R<sub>1</sub> = CH<sub>2</sub>OH, OMe, SMe; R = Et, R<sub>1</sub> = SMe), providing optically pure I by kinetic resolu. Compds. (.+.-)-I (R = R<sub>1</sub> = Me; R = Me, R<sub>1</sub> = CH<sub>2</sub>OH; R = Et, R<sub>1</sub> = SMe) have been resolved in multigram quantities using 0.003, 0.005, and 0.0004 mol % antibody catalysts, resp. Resolved compds. I are useful synthons for the construction of epothilones A-E and their analogs. Here, a formal synthesis of epothilone E (II), has been achieved starting from compd. I (R = Me, R<sub>1</sub> = CH<sub>2</sub>OH).

REFERENCE COUNT: 35 THERE ARE 35 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L36 ANSWER 10 OF 28 CAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 1999:78939 CAPLUS

DOCUMENT NUMBER: 130:209528

TITLE: A Highly Stereoselective Synthesis of Epothilone B

AUTHOR(S): White, James D.; Carter, Rich G.; Sundermann, Kurt F.

CORPORATE SOURCE: Department of Chemistry, Oregon State University, Corvallis, OR, OREGON, USA

SOURCE: Journal of Organic Chemistry (1999), 64(3), 684-685

CODEN: JOCEAH; ISSN: 0022-3263

PUBLISHER: American Chemical Society

DOCUMENT TYPE: Journal

LANGUAGE: English

OTHER SOURCE(S): CASREACT 130:209528

AB A convergent synthesis of epothilone B (I) that generates all seven of its asym. centers in a completely stereoselective fashion is described. The key step is the coupling of phosphonium salt II with aldehyde III.

REFERENCE COUNT: 27 THERE ARE 27 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L36 ANSWER 11 OF 28 CAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 1999:383492 CAPLUS

DOCUMENT NUMBER: 131:199535

TITLE: Total synthesis of epothilone E and related side-chain modified analogues via a Stille coupling based strategy

AUTHOR(S): Nicolaou, K. C.; King, N. P.; Finlay, M. R. V.; He, Y.; Roschangar, F.; Vourloumis, D.; Vallberg, H.; Sarabia, F.; Ninkovic, S.; Hepworth, D.

CORPORATE SOURCE: Department of Chemistry and The Skaggs Institute for Chemical Biology, The Scripps Research Institute, La

SOURCE: Jolla, CA, 92037, USA  
Bioorganic & Medicinal Chemistry (1999), 7(5), 665-697  
CODEN: BMECEP; ISSN: 0968-0896  
PUBLISHER: Elsevier Science Ltd.  
DOCUMENT TYPE: Journal  
LANGUAGE: English  
OTHER SOURCE(S): CASREACT 131:199535

AB A Stille coupling strategy has been utilized to complete a total synthesis of epothilone E from vinyl iodide I (R1 = I; R2 = H) and thiazolestannane II. The central core fragment I (R1 = I; R2 = H) and its trans-isomer III (R3 = I) were prepd. from triene IV (TBS = SiMe2CMe3) using ring-closing metathesis (RCM), and were subsequently coupled to a variety of alternative stannanes to provide a library of epothilone analogs I [R1 = 2-(5-acetoxypentyl)thiazol-4-yl, 2-(methylthio)thiazol-4-yl, 2-piperidinethiazol-4-yl, 2-methoxythiazol-4-yl, 2-ethoxythiazol-4-yl, thiazol-4-yl, thiazol-2-yl, thiazol-5-yl, 2-(hydroxymethyl)thiazol-4-yl, 2-(acetoxymethyl)thiazol-4-yl, 2-(fluoromethyl)thiazol-4-yl, 2-vinylthiazol-4-yl, 2-ethylthiazol-4-yl, 2-furyl, 2-thienyl, Ph,3-pyridyl, CH:COEtMe-(Z), R2 = H] and III [R3 = 2-(5-acetoxypentyl)thiazol-4-yl, 2-(methylthio)thiazol-4-yl, 2-piperidinethiazol-4-yl, 2-methoxythiazol-4-yl, 2-ethoxythiazol-4-yl, thiazol-4-yl, thiazol-2-yl, thiazol-5-yl, 2-(hydroxymethyl)thiazol-4-yl, 2-(acetoxymethyl)thiazol-4-yl, 2-(fluoromethyl)thiazol-4-yl, 2-vinylthiazol-4-yl, 2-ethylthiazol-4-yl, 2-furyl, 2-thienyl, Ph,3-pyridyl, CH:COEtMe-(Z)]. The Stille coupling approach was then used to prep. epothilone B analogs from the key macrolactone intermediate I (R1 = I, R2 = CH2OH) which was itself synthesized by a macrolactonization based strategy.

REFERENCE COUNT: 60 THERE ARE 60 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L36 ANSWER 12 OF 28 BIOSIS COPYRIGHT 2004 BIOLOGICAL ABSTRACTS INC. on STN  
ACCESSION NUMBER: 1999:182112 BIOSIS  
DOCUMENT NUMBER: PREV199900182112  
TITLE: **Desoxyepothilone B** (dEpoB): A tubulin stabilizing macrolide, is curative against human tumor xenografts that are refractory to taxol.  
AUTHOR(S): Zhang, X.-G.; Tan, Q.-H.; Chen, L. I.; Harris, C. R.; Kuduk, S. D.; Danishefsky, S. J.; Bertino, J. R.; Chou, T.-C.  
CORPORATE SOURCE: Mol. Pharmacol. Therap. Program, Memorial Sloan-Kettering Cancer Center, New York, NY 10021, USA  
SOURCE: Proceedings of the American Association for Cancer Research Annual Meeting, (March, 1999) Vol. 40, pp. 287. print.  
Meeting Info.: 90th Annual Meeting of the American Association for Cancer Research. Philadelphia, Pennsylvania, USA. April 10-14, 1999. American Association for Cancer Research.  
ISSN: 0197-016X.  
DOCUMENT TYPE: Conference; (Meeting)  
Conference; Abstract; (Meeting Abstract)  
LANGUAGE: English  
ENTRY DATE: Entered STN: 5 May 1999  
Last Updated on STN: 5 May 1999

L36 ANSWER 13 OF 28 CAPLUS COPYRIGHT 2004 ACS on STN  
ACCESSION NUMBER: 2000:739098 CAPLUS  
DOCUMENT NUMBER: 134:56491  
TITLE: **Epothilones**: microtubule stabilizing agents with enhanced activity against multidrug-resistant cell lines and tumors  
AUTHOR(S): Harris, Christina R.; Balog, Aaron; Savin, Kenneth; Danishefsky, Samuel J.; Chou, Ting Chao; Zhang, Xiu-Guo  
CORPORATE SOURCE: The Laboratory for Bioorganic Chemistry, New York, NY, 10021, USA  
SOURCE: Actualites de Chimie Therapeutique (1999), 25, 187-206  
CODEN: ACHTD9; ISSN: 0338-8999  
PUBLISHER: Editions Scientifiques et Medicales Elsevier  
DOCUMENT TYPE: Journal; General Review  
LANGUAGE: English

AB A review with 33 refs. on the synthesis and biol. activity of epothilones. Highly efficient, highly convergent total syntheses involving ring-closing olefin metathesis, Suzuki coupling, stereoselective aldol reactions and stereoselective Noyori redns. are discussed.

REFERENCE COUNT: 33 THERE ARE 33 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L36 ANSWER 14 OF 28 CAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 1999:322978 CAPLUS

DOCUMENT NUMBER: 131:124926

TITLE: A Unified and Quantitative Receptor Model for the Microtubule Binding of Paclitaxel and Epothilone

AUTHOR(S): Wang, Minmin; Xia, Xiaoyang; Kim, Yohan; Hwang, David; Jansen, Johanna M.; Botta, Maurizio; Liotta, Dennis C.; Snyder, James P.

CORPORATE SOURCE: Department of Chemistry, Emory University, Atlanta, GA, 30322, USA

SOURCE: Organic Letters (1999), 1(1), 43-46  
CODEN: ORLEF7; ISSN: 1523-7060

PUBLISHER: American Chemical Society

DOCUMENT TYPE: Journal

LANGUAGE: English

AB Paclitaxel and epothilone represent the two major classes of antimicrotubule agents that promote tubulin polymn. and, presumably, mitotic arrest during cell division. A common minireceptor binding site model at .beta.-tubulin has been constructed for these structurally divergent compds. Utilizing 20 amino acids identified in photoaffinity labeling expts., the 3-D model correlates measured and predicted Ki's with  $r = 0.99$  and  $\text{rms}(\text{.DELTA.Gcalc} - \text{.DELTA.Gexp}) = 0.2 \text{ kcal/mol}$ . In addn., the model predicts the affinity of compds. not used in the training set and explains much of the SAR for the paclitaxel and epothilone families.

REFERENCE COUNT: 52 THERE ARE 52 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L36 ANSWER 15 OF 28 CAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 1999:145094 CAPLUS

DOCUMENT NUMBER: 131:13461

TITLE: The microtubule-stabilizing agents epothilones A and B and their desoxy-derivatives induce mitotic arrest and apoptosis in human prostate cancer cells

AUTHOR(S): Sepp-Lorenzino, L.; Balog, A.; Su, D.-S.; Meng, D.; Timaul, N.; Scher, H. I.; Danishefsky, S. J.; Rosen, N.

CORPORATE SOURCE: Program in Cell Biology, Sloan-Kettering Institute for Cancer Research, New York, NY, 10021, USA

SOURCE: Prostate Cancer and Prostatic Diseases (1999), 2(1), 41-52  
CODEN: PCPDFW; ISSN: 1365-7852

PUBLISHER: Stockton Press

DOCUMENT TYPE: Journal

LANGUAGE: English

AB Epothilones are a new class of natural products that bind to tubulin and prevent the depolymn. of microtubules, although they have no structural similarity to paclitaxel. Taxanes are only marginally effective in the treatment of disseminated prostate cancer, although they may have useful activity when administered in combination with estramustine. Unlike paclitaxel, epothilones are not substrates for P-glycoprotein and are active in multidrug resistant cells. Epothilones A and B (EA, EB) have recently been synthesized in toto. In this report, we examine the effects of synthetic epothilones and their desoxy derivs., as well as paclitaxel, on prostate cancer cell lines. EB was the most active of these compds. in tissue culture ( $\text{IC}_{50}$ : 50-75 pM), four to ten-fold more potent than paclitaxel. EA and the desoxyderivatives of EA and EB (dEA, dEB) were also active, but less potent than EB. Each of these compds. causes mitotic block followed by apoptotic cell death. The relative potencies for cell cycle arrest and cytotoxicity directly correlate with the ability of the drugs to bind microtubules, stabilize mitotic spindles and induce the formation of interphase microtubule bundles. Therefore, synthetic epothilones are potent inhibitors of prostate cancer cell lines and work in a fashion similar to paclitaxel. Recently, we showed that farnesyl transferase inhibitors sensitize tumor cells to paclitaxel-induced mitotic

arrest. We now have extended these observations to show that paclitaxel and the epothilones synergize with FTI to arrest the growth of prostate cancer cells. Moreover, this occurs in DU145, a cell line that is not particularly sensitive to the FTI. The combination of FTI and epothilone represent a new potential clin. strategy for the treatment of advanced prostatic cancer.

REFERENCE COUNT: 42 THERE ARE 42 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L36 ANSWER 16 OF 28 CAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 1998:762086 CAPLUS  
DOCUMENT NUMBER: 129:343364  
TITLE: Methods for preparation of epothilone derivatives  
PATENT ASSIGNEE(S): Gesellschaft fuer Biotechnologische Forschung m.b.H.  
(GBF), Germany  
SOURCE: Ger. Offen., 2 pp.  
CODEN: GWXXBX  
DOCUMENT TYPE: Patent  
LANGUAGE: German  
FAMILY ACC. NUM. COUNT: 1  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
DE 19821954	A1	19981119	DE 1998-19821954	19980515
PRIORITY APPLN. INFO.:			DE 1997-19720250	19970515
OTHER SOURCE(S): MARPAT 129:343364				

AB Methods for prepn. of epothilone derivs. are characterized by: (a) proceeding from epothilones A, B, C or D, wherein the C(2)- and C(3)-atoms can be joined together through CH<sub>2</sub>CH(OH) or CH:CH and wherein one provides an (un)protected OH group at the resulting bond at C(3) and C(7); (b) oxidn. at C(16) to form a keto group; (c1) exchanging the oxygen of the keto-group to a :CH<sub>2</sub> group using Ph<sub>3</sub>P:CH<sub>2</sub>; and if necessary (d1) this :CH<sub>2</sub> group, with the help of the compd. RCH:CH<sub>2</sub>, is catalytically converted to a :CHR group [R = aliph. residue, (un)substituted Ph, heterocycle, esp. a pharmaceutically active residue]; or (c2) for the bond between C(16) and C(17) in known ways provides the CH:CH<sub>2</sub> group, and if necessary (d2) this group with the help of metathesis is converted into a :CHR group. Also claimed is the use of ozone to form the C(16) keto group. In addn., the reaction of the keto group with NaBH<sub>4</sub> followed by tosyl chloride and base or a Bamford-Stevens reaction to form the methylene compd. are claimed. Finally, rhodium, ruthenium, tungsten and molybdenum catalysts are claimed for the metathesis reactions.

L36 ANSWER 17 OF 28 MEDLINE on STN DUPLICATE 6

ACCESSION NUMBER: 1999080094 MEDLINE  
DOCUMENT NUMBER: PubMed ID: 9861050  
TITLE: **Desoxyepothilone B** is curative against human tumor xenografts that are refractory to paclitaxel.  
AUTHOR: Chou T C; Zhang X G; Harris C R; Kuduk S D; Balog A; Savin K A; Bertino J R; Danishefsky S J  
CORPORATE SOURCE: Molecular Pharmacology and Therapeutics Program, Sloan-Kettering Institute for Cancer Research, 1275 York Avenue, New York, NY 10021, USA.  
CONTRACT NUMBER: CA-08748 (NCI)  
CA-28824 (NCI)  
QM-18248  
SOURCE: Proceedings of the National Academy of Sciences of the United States of America, (1998 Dec 22) 95 (26) 15798-802.  
Journal code: 7505876. ISSN: 0027-8424.  
PUB. COUNTRY: United States  
DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)  
LANGUAGE: English  
FILE SEGMENT: Priority Journals  
ENTRY MONTH: 199901  
ENTRY DATE: Entered STN: 19990209  
Last Updated on STN: 19990209  
Entered Medline: 19990128

AB The epothilones are naturally occurring, cytotoxic macrolides that function through a paclitaxel (Taxol)-like mechanism. Although structurally dissimilar, both classes of molecules lead to the arrest of

cell division and eventual cell death by stabilizing cellular microtubule assemblies. The epothilones differ in their ability to retain activity against multidrug-resistant (MDR) cell lines and tumors where paclitaxel fails. In the current account, we focus on the relationship between epothilone and paclitaxel in the context of tumors with multiple drug resistance. The epothilone analogue Z-12,13-**desoxyepothilone B** (dEpoB) is >35,000-fold more potent than paclitaxel in inhibiting cell growth in the MDR DC-3F/ADX cell line. Various formulations, routes, and schedules of i.v. administration of dEpoB have been tested in nude mice. Slow infusion with a Cremophor-ethanol vehicle proved to be the most beneficial in increasing efficacy and decreasing toxicity. Although dEpoB performed similarly to paclitaxel in sensitive tumors xenografts (MX-1 human mammary and HT-29 colon tumor), its effects were clearly superior against MDR tumors. When dEpoB was administered to nude mice bearing our MDR human lymphoblastic T cell leukemia (CCRF-CEM/paclitaxel), dEpoB demonstrated a full curative effect. For human mammary adenocarcinoma MCF-7/Adr cells refractory to paclitaxel, dEpoB reduced the established tumors, markedly suppressed tumor growth, and surpassed other commonly used chemotherapy drugs such as adriamycin, vinblastine, and etoposide in beneficial effects.

L36 ANSWER 18 OF 28 LIFESCI COPYRIGHT 2004 CSA on STN  
 ACCESSION NUMBER: 1999:33776 LIFESCI  
 TITLE: A mAb recognizing a surface antigen of Mycobacterium tuberculosis enhances host survival  
 AUTHOR: Teitelbaum, R.; Freedman, A.G.; Chen, B.; Robbins, J.B.; Unanue, E.; Casadevall, A.; Bloom, B.R.  
 CORPORATE SOURCE: Department of Microbiology and Immunology of the Albert Einstein College of Medicine 1300 Morris Park Avenue, Bronx, NY 10064 USA; E-mail: bloom@aecom.yu.edu  
 SOURCE: Proceedings of the National Academy of Sciences, USA [Proc. Natl. Acad. Sci. USA], (19981222) vol. 95, no. 26, pp. 15688-15693.  
 ISSN: 0027-8424.  
 DOCUMENT TYPE: Journal  
 FILE SEGMENT: F; J  
 LANGUAGE: English  
 SUMMARY LANGUAGE: English

AB Murine mAbs reactive with the surface of Mycobacterium tuberculosis were assayed for their ability to affect the course of infection in mice challenged with virulent organisms. An IgG3 mAb (9d8) specific for arabinomannan and reactive with purified antigen from a clinical isolate of M. tuberculosis conferred partial protection on mice after respiratory challenge (30-60% survival >75 days; P=0.05). Control mice pretreated with an irrelevant mAb of the same isotype succumbed to tuberculosis within 30 days. Mice with gene disruptions in interferon gamma and major histocompatibility complex Class II also were partially protected from challenge. The protective mAb was neither bactericidal nor inhibitory of infection or bacterial replication. Nevertheless, it profoundly altered the nature of the granulomas in the infected lungs. Mice treated with mAb 9d8 and challenged with M. tuberculosis localized the pathogen within granuloma centers suggesting that the mAb conferred protection by enhancing a cellular immune response.

L36 ANSWER 19 OF 28 MEDLINE on STN DUPLICATE 7  
 ACCESSION NUMBER: 1998356211 MEDLINE  
 DOCUMENT NUMBER: PubMed ID: 9689134  
 TITLE: **Desoxyepothilone B**: an efficacious microtubule-targeted antitumor agent with a promising in vivo profile relative to epothilone B.  
 AUTHOR: Chou T C; Zhang X G; Balog A; Su D S; Meng D; Savin K; Bertino J R; Danishefsky S J  
 CORPORATE SOURCE: Molecular Pharmacology and Therapeutics Program, 1275 York Avenue, New York, NY 10021, USA.  
 CONTRACT NUMBER: CA-28824 (NCI)  
 CA-GM 72231 (NCI)  
 SOURCE: Proceedings of the National Academy of Sciences of the United States of America, (1998 Aug 4) 95 (16) 9642-7.  
 Journal code: 7505876. ISSN: 0027-8424.  
 PUB. COUNTRY: United States  
 DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English  
FILE SEGMENT: Priority Journals  
ENTRY MONTH: 199809  
ENTRY DATE: Entered STN: 19980917  
Last Updated on STN: 19980917  
Entered Medline: 19980908

AB A new class of 16-membered macrolides, the epothilones (Epos), has been synthesized and evaluated for antitumor potential in vitro and in vivo. Recent studies in these and other laboratories showed that epothilones and paclitaxel (paclitaxel) share similar mechanisms of action in stabilizing microtubule arrays as indicated by binding-displacement studies, substitution for paclitaxel in paclitaxel-dependent cell growth, and electron microscopic examinations. The present study examined cell growth-inhibitory effects in two rodent and three human tumor cell lines and their drug-resistant sublines. Although paclitaxel showed as much as 1, 970-fold cross-resistance to the sublines resistant to paclitaxel, adriamycin, vinblastine, or actinomycin D, most epothilones exhibit little or no cross-resistance. In multidrug-resistant CCRF-CEM/VBL100 cells, IC50 values for EpoA (1), EpoB (2), desoxyEpoA (3) (dEpoA), desoxyEpoB (4) (dEpoB), and paclitaxel were 0.02, 0.002, 0.012, 0.017, and 4.14 microM, respectively. In vivo studies, using i.p. administration, indicated that the parent, EpoB, was highly toxic to mice and showed little therapeutic effect when compared with a lead compound, dEpoB. More significantly, dEpoB (25-40 mg/kg, Q2Dx5, i.p.) showed far superior therapeutic effects and lower toxicity than paclitaxel, doxorubicin, camptothecin, or vinblastine (at maximal tolerated doses) in parallel experiments. For mammary adenocarcinoma xenografts resistant to adriamycin, MCF-7/Adr, superior therapeutic effects were obtained with dEpoB compared with paclitaxel when i.p. regimens were used. For ovarian adenocarcinoma xenografts, SK-OV-3, dEpoB (i.p.), and paclitaxel (i. v.) gave similar therapeutic effects. In nude mice bearing a human mammary carcinoma xenograft (MX-1), marked tumor regression and cures were obtained with dEpoB.

L36 ANSWER 20 OF 28 CAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 1998:732784 CAPLUS  
DOCUMENT NUMBER: 130:81320  
TITLE: Easy access to the epothilone family - synthesis of epothilone B  
AUTHOR(S): Mulzer, Johann; Mantoulidis, Andreas; Ohler, Elisabeth  
CORPORATE SOURCE: Inst. fur Organische Chemie, Univ. Wien, Vienna, A-1090, Austria  
SOURCE: Tetrahedron Letters (1998), 39(47), 8633-8636  
CODEN: TELEAY; ISSN: 0040-4039  
PUBLISHER: Elsevier Science Ltd.  
DOCUMENT TYPE: Journal  
LANGUAGE: English  
OTHER SOURCE(S): CASREACT 130:81320

AB An easy access to four out of five naturally occurring epothilones (A-E) is reported. Key steps are an enantioselective Mukaiyama type aldol reaction, (E)- and (Z)-selective olefinations, and a sulfone alkylation.  
REFERENCE COUNT: 29 THERE ARE 29 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L36 ANSWER 21 OF 28 MEDLINE on STN DUPLICATE 8

ACCESSION NUMBER: 1998132596 MEDLINE  
DOCUMENT NUMBER: PubMed ID: 9465021  
TITLE: Farnesyl transferase inhibitors cause enhanced mitotic sensitivity to taxol and epothilones.  
AUTHOR: Moasser M M; Sepp-Lorenzino L; Kohl N E; Oliff A; Balog A; Su D S; Danishefsky S J; Rosen N  
CORPORATE SOURCE: Department of Medicine, Sloan-Kettering Institute, Memorial Sloan-Kettering Cancer Center, 1275 York Avenue, New York, NY 10021, USA.  
CONTRACT NUMBER: P50CA68425-02 (NCI)  
SOURCE: Proceedings of the National Academy of Sciences of the United States of America, (1998 Feb 17) 95 (4) 1369-74. Journal code: 7505876. ISSN: 0027-8424.  
PUB. COUNTRY: United States  
DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)  
LANGUAGE: English

FILE SEGMENT: Priority Journals  
ENTRY MONTH: 199803  
ENTRY DATE: Entered STN: 19980326  
Last Updated on STN: 19980326  
Entered Medline: 19980319

AB An important class of cellular proteins, which includes members of the p21ras family, undergoes posttranslational farnesylation, a modification required for their partition to membranes. Specific farnesyl transferase inhibitors (FTIs) have been developed that selectively inhibit the processing of these proteins. FTIs have been shown to be potent inhibitors of tumor cell growth in cell culture and in murine models and at doses that cause little toxicity to the animal. These data suggest that these drugs might be useful therapeutic agents. We now report that, when FTI is combined with some cytotoxic antineoplastic drugs, the effects on tumor cells are additive. No interference is noted. Furthermore, FTI and agents that prevent microtubule depolymerization, such as taxol or epothilones, act synergistically to inhibit cell growth. FTI causes increased sensitivity to induction of metaphase block by these agents, suggesting that a farnesylated protein may regulate the mitotic check point. The findings imply that FTI may be a useful agent for the treatment of tumors with wild-type ras that are sensitive to taxanes.

L36 ANSWER 22 OF 28 CAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 1998:150476 CAPLUS

DOCUMENT NUMBER: 128:230166

TITLE: Total synthesis of epothilone E and analogs with modified side chains through the Stille coupling reaction

AUTHOR(S): Nicolaou, K. C.; He, Yun; Roschangar, Frank; King, N. Paul; Vourloumis, Dionisios; Li, Tianhu

CORPORATE SOURCE: Department of Chemistry, Skaggs Inst. for Chemical Biology, Scripps Res. Inst., La Jolla, CA, 92037, USA

SOURCE: Angewandte Chemie, International Edition (1998), 37(1/2), 84-87

CODEN: ACIEF5; ISSN: 1433-7851

PUBLISHER: Wiley-VCH Verlag GmbH

DOCUMENT TYPE: Journal

LANGUAGE: English

OTHER SOURCE(S): CASREACT 128:230166

AB The first total synthesis of epothilone E [I; R = 2-(hydroxymethyl)thiazol-4-yl, X = O] in which an olefin metathesis is used to form the macrocyclic lactone and a Stille coupling reaction is used to form the side chain is reported. The Stille coupling reaction was used to prep. deoxygenated side-chain analogs I [R = thiazol-4-yl, thiazol-5-yl, thiazol-2-yl, 2-(5-acetoxypentyl)thiazol-4-yl, 2-piperidinethiazol-4-yl, 2-(methylthio)thiazol-4-yl, 2-furyl, 2-thienyl, Ph, 3-pyridyl; X = bond].

REFERENCE COUNT: 55 THERE ARE 55 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L36 ANSWER 23 OF 28 CAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 1997:665094 CAPLUS

DOCUMENT NUMBER: 127:293040

TITLE: Total Syntheses of Epothilones A and B

AUTHOR(S): Meng, Dongfang; Bertinato, Peter; Balog, Aaron; Su, Dai-Shi; Kamenecka, Ted; Sorensen, Erik; Danishefsky, Samuel J.

CORPORATE SOURCE: Laboratory for Bioorganic Chemistry, Sloan-Kettering Institute for Cancer Research, New York, NY, 10021, USA

SOURCE: Journal of the American Chemical Society (1997), 119(42), 10073-10092

CODEN: JACSAT; ISSN: 0002-7863

PUBLISHER: American Chemical Society

DOCUMENT TYPE: Journal

LANGUAGE: English

OTHER SOURCE(S): CASREACT 127:293040

AB Convergent, stereocontrolled total syntheses of the microtubule-stabilizing macrolides epothilones A (I; R = H) and B (I; R = Me) have been achieved. Four distinct ring-forming strategies were pursued. Of these four, three were reduced to practice. In one approach, the action of a base on a substance possessing an acetate ester and a nonenolizable

aldehyde brought about a remarkably effective macroaldolization simultaneously creating the C2-C3 bond and the hydroxyl-bearing stereocenter at C-3. Alternatively, the 16-membered macrolide of the epothilones could be fashioned through a C12-C13 ring-closing olefin metathesis and through macrolactonization of the appropriate hydroxy acid. The application of a stereospecific B-alkyl Suzuki coupling strategy permitted the establishment of a cis C12-C13 olefin, thus setting the stage for an eventual site- and diastereoselective epoxidn. reaction. The development of a novel cyclopropane solvolysis strategy for incorporating the geminal Me groups of the epothilones, and the use of Lewis acid catalyzed diene-aldehyde cyclocondensation (LACDAC) and asym. allylation methodol. are also noteworthy.

L36 ANSWER 24 OF 28 CAPLUS COPYRIGHT 2004 ACS on STN  
 ACCESSION NUMBER: 1997:430309 CAPLUS  
 DOCUMENT NUMBER: 127:108793  
 TITLE: Stereoselective syntheses and evaluation of compounds in the 8-desmethylepothilone A series: some surprising observations regarding their chemical and biological properties  
 AUTHOR(S): Balog, Aaron; Betinato, Peter; Su, Dai-Shi; Meng, Dongfang; Sorensen, Erik; Danishefsky, Samuel J.; Zheng, Yu-Huang; Chou, Ting-Chao; He, Lifeng; Horwitz, Susan B.  
 CORPORATE SOURCE: Lab. Bioorganic Chem., Sloan-Kettering Inst. Cancer Res., New York, NY, 10021, USA  
 SOURCE: Tetrahedron Letters (1997), 38(26), 4529-4532  
 CODEN: TELEAY; ISSN: 0040-4039  
 PUBLISHER: Elsevier  
 DOCUMENT TYPE: Journal  
 LANGUAGE: English  
 OTHER SOURCE(S): CASREACT 127:108793  
 AB The title compds. have been synthesized in a convergent way by recourse to a Weiler type dianion construction.  
 REFERENCE COUNT: 18 THERE ARE 18 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L36 ANSWER 25 OF 28 CAPLUS COPYRIGHT 2004 ACS on STN  
 ACCESSION NUMBER: 1997:175662 CAPLUS  
 DOCUMENT NUMBER: 126:225133  
 TITLE: Remote Effects in Macrolide Formation through Ring-Forming Olefin Metathesis: An Application to the Synthesis of Fully Active Epothilone Congeners  
 AUTHOR(S): Meng, Dongfang; Su, Dai-Shi; Balog, Aaron; Bertinato, Peter; Sorensen, Erik J.; Danishefsky, Samuel J.; Zheng, Yu-Huang; Chou, Ting-Chao; He, Lifeng; Horwitz, Susan B.  
 CORPORATE SOURCE: Laboratories for Bioorganic Chemistry and Biochemical Pharmacology, Sloan-Kettering Institute for Cancer Research, New York, NY, 10021, USA  
 SOURCE: Journal of the American Chemical Society (1997), 119(11), 2733-2734  
 CODEN: JACSAT; ISSN: 0002-7863  
 PUBLISHER: American Chemical Society  
 DOCUMENT TYPE: Journal  
 LANGUAGE: English  
 OTHER SOURCE(S): CASREACT 126:225133  
 AB A ring closing olefin metathesis strategy for the synthesis of the previously encountered **desoxyepothilone A** (I) is described. A merging of the alkyl segment II (carbons 12-21) and acyl segment III (carbons 3-11) through an intermol. aldol-condensation reaction provided substrates needed for ring closing olefin metathesis. Thus, thiazole IV underwent olefin metathesis in C6H6 contg. 50 mol % (PhCH:)[P(cyclohexyl)3]2RuCl2 to give 65% II and its E-isomer (Z:E 1:2). The results of these cyclization indicate a remarkable sensitivity to permutations of functionality at centers remote from the site of olefin metathesis. The in vitro biol. activity of E and Z **desoxyepothilone** as well as several related congeners is also described. I has IC50 range of 0.012-0.022 .mu.M against drug-sensitive and -resistant human leukemic CCRF-CEM cell lines.



L36 ANSWER 26 OF 28 CAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 1997:714314 CAPLUS  
DOCUMENT NUMBER: 127:358730  
TITLE: Structure-activity relationships of the epothilones and the first in vivo comparison with paclitaxel  
AUTHOR(S): Su, Dai-Shi; Balog, Aaron; Meng, Dongfang; Bertinato, Peter; Danishefsky, Samuel J.; Zheng, Yu-Huang; Chou, Ting-Chao; He, Lifeng; Horwitz, Susan B.  
CORPORATE SOURCE: Laboratory for Bioorganic Chemistry, Sloan-Kettering Institute for Cancer Research, New York, NY, 10021, USA  
SOURCE: Angewandte Chemie, International Edition in English (1997), 36(19), 2093-2096  
CODEN: ACIEAY; ISSN: 0570-0833  
PUBLISHER: Wiley-VCH  
DOCUMENT TYPE: Journal  
LANGUAGE: English  
AB The structure-activity relationships of the epothilones and 18 derivs. and analogs were studied. An in vivo comparison of the chemotherapeutic effect of epothilone B with that of paclitaxel was also studied. The chemotherapeutic effect of daily doses of epothilone B (0.7 mg/kg) and paclitaxel (2 mg/kg) in CB-17 SCID mice bearing drug-resistant human CCRF-CEM/VBL xenografts were T/C = 0.33 and T/C = 0.70, resp.  
REFERENCE COUNT: 34 THERE ARE 34 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L36 ANSWER 27 OF 28 CAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 1998:729 CAPLUS  
DOCUMENT NUMBER: 128:88685  
TITLE: Metathesis vs metastasis: the chemistry and biology of the epothilones  
AUTHOR(S): Finlay, Ray  
CORPORATE SOURCE: Dep. Chemistry, The Skaggs Inst. for Chemical Biol., The Scripps Res. Inst., La Jolla, CA, 92037, USA  
SOURCE: Chemistry & Industry (London) (1997), (24), 991-996  
CODEN: CHINAG; ISSN: 0009-3068  
PUBLISHER: Society of Chemical Industry  
DOCUMENT TYPE: Journal; General Review  
LANGUAGE: English  
AB A review with 15 refs. on a recent entry onto the scene of potentially useful natural products, the epothilones A - E, providing valuable information for the fight against cancer via their interaction with microtubules.  
REFERENCE COUNT: 26 THERE ARE 26 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L36 ANSWER 28 OF 28 CAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 1997:302059 CAPLUS  
DOCUMENT NUMBER: 127:4948  
TITLE: Total synthesis of (-)-epothilone B: an extension of the Suzuki coupling method and insights into structure-activity relationships of the epothilones  
AUTHOR(S): Su, Dai-Shi; Meng, Dongfang; Bertinato, Peter; Balog, Aaron; Sorensen, Erik J.; Danishefsky, Samuel J.; Zheng, Yu-Huang; Chou, Ting-Chao; He, Lifeng; Horwitz, Susan B.  
SOURCE: Angewandte Chemie, International Edition in English (1997), 36(7), 757-759  
CODEN: ACIEAY; ISSN: 0570-0833  
PUBLISHER: VCH  
DOCUMENT TYPE: Journal  
LANGUAGE: English  
OTHER SOURCE(S): CASREACT 127:4948  
AB (-)-Epothilone B (I; R = Me, X = O) and **desoxyepothilone B** (I; R = Me, X = bond) were prepd. via Suzuki coupling of (Z)-vinyl iodide II with borane III. I (R = H, Me, X = O, bond) and the E-isomers of I (R = H, Me, X = bond) were tested for efficacy against drug-sensitive and resistant CCRF-CEM cell lines (IC50 = 0.0004 - 0.262 .mu.M).  
REFERENCE COUNT: 27 THERE ARE 27 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

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